U.S. DEPARTMENT OF COMMERCI Patent and Trademark Office

SEARCH REQUEST FORM 58432

| Requestor's Name: Nata (12) | Serial Number: $09/$ | 867034 |
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| Search Topic: Please write a detailed statement of search topic. De that may have a special meaning. Give examples or a copy of the sequence. You may include a copy o | scribe specifically as possible the subject matter relevant citations, authors keywords, etc., if kn of the broadest and/or most relevant claim(s). | r to be searched. Define any terms own. For sequences, please attach |
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| Number of Databases: | Structure | DARC/Questel |
| | Bibliographic | Other |

TO-1590 (9-90)

USCOMM-DC 90-3952

(FILE *CAPLUS* ENTERED AT 10:04:19 ON 18 JAN 2002) 112953 SEA FILE=CAPLUS ABB=ON PLU=ON ((COLON OR COLONIC)(S)(CA L1 NCER? OR CARCIN? OR TUMOUR OR TUMOR OR NEOPLAS?)) OR METAST? 20 SEA FILE=CAPLUS ABB=ON PLU=ON L1 AND (CSG OR COLON L2 SPECIF? GENE) ANSWER 1 OF 20 CAPLUS COPYRIGHT 2002 ACS L22002:10740 CAPLUS ACCESSION NUMBER: Use of colon specific TITLE: gene polypeptides in diagnosing, monitoring, staging, imaging and treating colon cancer Macina, Roberto A.; Pillai, Rajeswari INVENTOR(S): Diadexus, Inc., USA PATENT ASSIGNEE(S): PCT Int. Appl., 135 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent English LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: APPLICATION NO. PATENT NO. KIND DATE DATE ----_____ _____ _____ A2 20020103 WO 2001-US20724 20010628 WO 2002000939 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, ΤG P 20000628 US 2000-214515 PRIORITY APPLN. INFO.: The invention relates to colon specific gene (CSG) polypeptides, polynucleotides encoding the polypeptides, methods for producing the polypeptides, in particular by expressing the polynucleotides, and agonists and antagonists of the polypeptides. The present invention includes methods of diagnosing metastases or staging of colon cancer in a patient by comparing CSG

particular by expressing the polynucleotides, and agonists and antagonists of the polypeptides. The present invention includes methods of diagnosing metastases or staging of colon cancer in a patient by comparing CSG expression levels in cells, tissues and body fluids of colon cancer patients and normal human control. Increased expression of CSG indicates progressive cancer while decreased CSG expression is correlated with cancer that is regressing or in remission. The invention further relates to methods for utilizing such polynucleotides, polypeptides, agonists and antagonists for applications, which relate, in part, to research, diagnostic and clin. arts. Antibodies to CSG polypeptides can be labeled for detection in tissues which would be useful in detecting colon cancer via imaging and therapy. Vaccines contg. CSG proteins are another embodiment of the invention.

L2 ANSWER 2 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2001:886514 CAPLUS

DOCUMENT NUMBER:

136:34276

TITLE:

Method of diagnosing, monitoring, staging,

imaging and treating colon

cancer

INVENTOR(S):

Macina, Roberto A.; Chen, Sei-yu; Pluta, Jason;

Sun, Yongming; Recipon, Herve

PATENT ASSIGNEE(S):

Diadexus, Inc., USA

SOURCE:

PCT Int. Appl., 116 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PA' | TENT | NO. | | KI | ND | DATE | | | A. | PPLI | CATI | ON N | ο. | DATE | | |
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| WO | 2001 | 0925 | 28 | A | 2 | 2001 | 1206 | | W | O 20 | 01-U | S175 | 83 | 2001 | 0529 | |
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| | • | CN, | CO, | CR, | CU, | CZ, | DE, | DK, | DM, | DZ, | EE, | ES, | FI, | GB, | GD, | GE, |
| | | GH, | GM, | HR, | HU, | ID, | IL, | IN, | IS, | JP, | ΚE, | KG, | KP, | KR, | ΚZ, | LC, |
| | | LK, | LR, | LS, | LT, | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | ΜZ, | NO, |
| | | ΝZ, | PL, | PT, | RO, | RU, | SD, | SE, | SG, | SI, | SK, | SL, | ТJ, | TM, | TR, | TT, |
| | | TZ, | UA, | UG, | US, | UZ, | VN, | YU, | ZA, | ZW, | AM, | ΑZ, | BY, | KG, | ΚZ, | MD, |
| | | RU, | ТJ, | TM | | | | | | | | | | • | | |
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| | | CY, | DE, | DK, | ES, | FI, | FR, | GB, | GR, | ΙE, | IT, | LU, | MC, | NL, | PT, | SE, |
| | | TR, | BF, | ВJ, | CF, | CG, | CI, | CM, | GA, | GN, | GW, | ML, | MR, | ΝE, | SN, | TD, |
| | | ma | | | | | | | | | | | | | | |

PRIORITY APPLN. INFO.:

US 2000-207383 P 20000526

The invention relates to CSG (colon-

specific genes) polypeptides, polynucleotides

encoding the polypeptides, methods for producing the polypeptides, in particular by expressing the polynucleotides, and agonists and antagonists of the polypeptides. The invention further relates to methods for utilizing such polynucleotides, polypeptides, agonists and antagonists for applications, which relate, in part, to research, diagnostic and clin. arts.

ANSWER 3 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2001:730999 CAPLUS

DOCUMENT NUMBER:

135:284064

TITLE:

Colon cancer-associated cDNA

sequences and methods for diagnosing, monitoring, staging, imaging and treating

colon cancers

INVENTOR(S):

Yang, Fei; Piderit, Alejandra; Hu, Ping;

Recipon, Herve; Macina, Roberto A.

PATENT ASSIGNEE(S): SOURCE:

Diadexus, Inc., USA PCT Int. Appl., 105 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE PATENT NO. KIND DATE

> Shears 308-4994 Searcher :

WO 2001073030 A2 20011004 WO 2001-US9737 20010326

W: AU, CA, JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,

NL, PT, SE, TR

PRIORITY APPLN. INFO.: US 2000-192667 P 20000328

AB The present invention provides fifty seven cDNA fragment sequence which are diagnostic markers for colon cancer.

In addn., antibodies immunospecific for these markers are provided. Vectors, hosts cells and methods for producing these markers, as well as methods and tools for using these markers in detecting,

diagnosing, monitoring, staging, prognosticating, imaging and treating colon cancer are also provided.

L2 ANSWER 4 OF 20 CAPLUS COPYRIGHT 2002 ACS

TITLE: hTERT expression and cellular immunity in

2001:534845 CAPLUS

gastric cancer and precancerosis

AUTHOR(S): Yao, Xixian; Yin, Lei; Zhang, Jieying; Bai,

Wenyuan; Li, Yingmin; Sun, Zhongcheng

CORPORATE SOURCE: The Department of Digestive Medicine, the 2nd

Hospital, Hebei Medical University, Shijiazhuang, 050000, Peop. Rep. China

SOURCE: Shijie Huaren Xiaohua Zazhi (2001), 9(5),

508-512

CODEN: SHXZF2

PUBLISHER: Shijie Weichangbingxue Zazhishe

DOCUMENT TYPE: Journal LANGUAGE: Chinese

ACCESSION NUMBER:

The expression of human telomerase reverse transcriptase in gastric carcinomas and precancerous lesions were studied for evaluating the immune state of such patients and the clin. implications of hTERT and immune state for the diagnosis, treatment and prognosis of gastric cancer. In situ hybridization was used to detect the expression of hTERT mRNA in 116 endoscopic biopsies of gastric mucosa. Tissue samples were analyzed as follows: 30 cases of chronic superficial gastritis(CSG), 44 of precancerous lesions (including 27 of chronic atrophy gastritis, 8 of hyperplastic ploy and 9 of gastric ulcer) and 42 of gastric cancer (GC). At mean time, the T lymphocyte subsets(CD3+, CD4+/CD8+) and natural killer cell(NK) in peripheral blood were detd. by flow cytometrical anal. (FCM) in 30 cases of CSG, 27 of precancerosis (chronic atrophy gastritis, CAG), 42 of GC and the data were compared with those of normal controls (NC). The pos. rate of hTERT varied as follows: 0% (0/30) in CSG, 36% (16/44) in precancerous lesions and 86% (36/42) in GC. expression of hTERT mRNA was not assocd. with patients gender, tumor location, macroscopic type, lymph node metastasis and degree of differentiation. The CD3+ and CD4+ of CSG were lower than that of NC (P < 0.05). Meanwhile, the T lymphocyte subsets (CD3+, CD4+, CD4+/CD8+ ratio) were remarkably lower than that of NC and CSG . Furthermore with the tumor progression, the function of T cells was weakened gradually. expression of telomerase may be a crucial step in gastric carcinogenesis and increased hTERT mRNA may serve as a novel marker for diagnosis of gastric cancer. The immune state of patients with gastric cancer and precancerosis was somewhat depressed, which indicates the importance of cellular immunity in cancer patients.

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ANSWER 5 OF 20 CAPLUS COPYRIGHT 2002 ACS
                        2001:475327 CAPLUS
ACCESSION NUMBER:
                        135:207449
DOCUMENT NUMBER:
                        Nucleic acid-based ribozyme and DNAzyme
TITLE:
                        modulators of gene expression
                        McSwiggen, James; Usman, Nassim; Blatt,
INVENTOR(S):
                        Lawrence; Beigelman, Leonid; Burgin, Alex;
                         Karpeisky, Alexander; Matulic-Adamic, Jasenka;
                         Sweedler, David; Draper, Kenneth; Chowrira,
                         Bharat; Stinchcomb, Dan; Beaudry, Amber; Zinnen,
                         Shawn; Lugwig, Janos; Sproat, Brian S.
                         Ribozyme Pharmaceuticals, Inc., USA
PATENT ASSIGNEE(S):
SOURCE:
                         PCT Int. Appl., 717 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
PATENT INFORMATION:
                                          APPLICATION NO. DATE
    PATENT NO.
                     KIND DATE
                     ____
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                           20010308 WO 2000-US23998 20000830
    WO 2001016312 A2
       AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
        CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
        HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
        LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ
    RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA,
        GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                          US 1999-PV151713 19990831
                                                          19990927
                                          US 1999-406643
                                          US 1999-PV156467 19990927
                                          US 1999-PV156236 19990927
                                          US 1999-436430
                                                          19991108
                                          US 1999-PV169100 19991206
                                          US 1999-PV173612 19991229
                                          US 1999-474432
                                                          19991229
                                          US 1999-476387
                                                           19991230
                                          US 2000-498824
                                                           20000204
                                          US 2000-531025
                                                           20000320
                                          US 2000-PV197769 20000414
                                          US 2000-578223
                                                           20000523
    Novel nucleic acid mols. useful as inhibitors of gene expression,
AB
    compns., and methods for their use are provided. The invention
    features novel nucleic acid-based techniques (e.g., enzymic nucleic
    acid mols. (ribozymes), antisense nucleic acids, 2-5A antisense
    chimeras, triplex DNA, and antisense nucleic acids contg.
    RNA-cleaving chem. groups) and their use to modulate the expression
    of mol. targets impacting the development and progression of
    cancers, diabetes, obesity, Alzheimer's disease diseases,
    age-related diseases, and/or hepatitis B infections and related
    conditions. Catalytic nucleic acids were designed for site-specific
    cleavage of human mRNA targets encoding protein tyrosine phosphatase
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1b, methionine aminopeptidase, .beta.-secretase, presenilin-1,

telomerase, and hepatitis B virus genes. Methods for chem. synthesis of modified nucleoside triphosphates (NTPs) and RNA polymerase-catalyzed incorporation of modified NTPs into catalytic oligonucleotides are also provided. [This abstr. record os one of 6 records for this document necessitated by the large no. of index

epidermal growth factor receptor-2 (HER2/c-erb2/neu), phospholamban,

entries required to fully index the document and publication system constraints.].

L2 ANSWER 6 OF 20 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:416790 CAPLUS

DOCUMENT NUMBER:

135:30984

TITLE:

Cancer-specific gene products for diagnosing, monitoring, staging, imaging and treating

prostate cancer

INVENTOR(S):

Ali, Shujath; Cafferkey, Robert; Recipon, Herve;

Sun, Yongming

PATENT ASSIGNEE(S): SOURCE:

Diadexus, Inc., USA PCT Int. Appl., 52 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2001039798 A1 20010607 WO 2000-US32927 20001205

W: CA, JP

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR

PRIORITY APPLN. INFO.:

US 1999-169083 P 19991206

AB The present invention provides new markers and methods for detecting, diagnosing, monitoring, staging, prognosticating, imaging and treating prostate cancer. The markers are cancer-specific gene products.

REFERENCE COUNT:

6

REFERENCE(S):

- (1) Adams, M; Nature 1995, V377, P3 CAPLUS
- (2) Bussemakers, M; European Urology 1999, V35(5-6), P408 CAPLUS
- (3) Hoon; Journal of Immunology 1995, V154, P730 CAPLUS
- (4) Human Genome Sci Inc; WO 9639435 A1 1996 CAPLUS
- (5) Raming; Receptors and Channels 1998, V6(2), P141 CAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 7 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

2001:400023 CAPLUS

Correction of: 2001:294219

DOCUMENT NUMBER:

135:16022

Correction of: 134:337614

Nucleic acid-based ribozyme and DNAzyme modulators of gene expression

INVENTOR(S): McSwiggen, James; Usman, Nassim; Blatt,

Lawrence; Beigelman, Leonid; Burgin, Alex; Karpeisky, Alexander; Matulic-adamic, Jasenka; Sweedler, David; Draper, Kenneth; Chowrira, Bharat; Stinchcomb, Dan; Beaudry, Amber; Zinnen,

Shawn; Lugwig, Janos; Sproat, Brian S.

PATENT ASSIGNEE(S):

Ribozyme Pharmaceuticals, Inc., USA

SOURCE:

TITLE:

PCT Int. Appl., 717 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:
PATENT INFORMATION:

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KIND DATE
                                             APPLICATION NO.
                                                               DATE
     PATENT NO.
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                                             WO 2000-US23998 20000830
                             20010308
     WO 2001016312 A2
         AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
         CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
         HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
         LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ
     RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG
                                             US 1999-PV151713 19990831
PRIORITY APPLN. INFO.:
                                             US 1999-406643
                                                              19990927
                                             US 1999-PV156467 19990927
                                             US 1999-PV156236 19990927
                                             US 1999-436430
                                                               19991108
                                             US 1999-PV169100 19991206
                                             US 1999-PV173612 19991229
                                             US 1999-474432
                                                               19991229
                                                               19991230
                                             US 1999-476387
                                             US 2000-498824
                                                               20000204
                                             US 2000-531025
                                                               20000320
                                             US 2000-PV197769 20000414
                                             US 2000-578223
                                                               20000523
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Novel nucleic acid mols. useful as inhibitors of gene expression, AB compns., and methods for their use are provided. The invention features novel nucleic acid-based techniques (e.g., enzymic nucleic acid mols. (ribozymes), antisense nucleic acids, 2-5A antisense chimeras, triplex DNA, and antisense nucleic acids contg. RNA-cleaving chem. groups) and their use to modulate the expression of mol. targets impacting the development and progression of cancers, diabetes, obesity, Alzheimer's disease diseases, age-related diseases, and/or hepatitis B infections and related conditions. Catalytic nucleic acids were designed for site-specific cleavage of human mRNA targets encoding protein tyrosine phosphatase 1b, methionine aminopeptidase, .beta.-secretase, presenilin-1, epidermal growth factor receptor-2 (HER2/c-erb2/neu), phospholamban, telomerase, and hepatitis B virus genes. Methods for chem. synthesis of modified nucleoside triphosphates (NTPs) and RNA polymerase-catalyzed incorporation of modified NTPs into catalytic oligonucleotides are also provided. [This abstr. record os one of 6 records for this document necessitated by the large no. of index entries required to fully index the document and publication system constraints.].

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L2 ANSWER 8 OF 20 CAPLUS COPYRIGHT 2002 ACS
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ACCESSION NUMBER: 2001:294219 CAPLUS

Correction of: 2001:168136

DOCUMENT NUMBER:

134:337614

Correction of: 134:233606

TITLE:

Nucleic acid-based ribozyme and DNAzyme

modulators of gene expression

INVENTOR(S):

McSwiggen, James; Usman, Nassim; Blatt, Lawrence; Beigelman, Leonid; Burgin, Alex; Karpeisky, Alexander; Matulic-adamic, Jasenka; Sweedler, David; Draper, Kenneth; Chowrira,

Bharat; Stinchcomb, Dan; Beaudry, Amber; Zinnen,

Shawn; Lugwig, Janos; Sproat, Brian S.

PATENT ASSIGNEE(S): Ribozyme Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 717 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE: PATENT INFORMATION:

| I | PATENT NO. KIND DATE | | | | APPLICATION NO. DATE | | | | | | | | | | | | |
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| | | HU, | ID, | IL, | IN, | IS, | JP, | KE, | KG, | KP, | KR, | KZ, | LC, | LK, | LR, | LS, | LT, |
| | | LU, | LV, | MA, | MD, | MG, | MK, | MN, | MW, | MX, | MZ, | NO, | NZ | | | | |
| F | RW: | | | | | | | | | | | | | ES, | FI, | FR, | GΑ, |
| | | | | | | | | | | | | | | | TD, | | |
| PRIOR | ITY | APPI | LN. | INFO | . : | - | • | • | | U | S 19 | 99-P | V151 | 713 | 19990 | 0831 | |
| | | | | | | | | | | U | S 19 | 99-4 | 0664 | 3 | 19990 | 0927 | |
| | | | | | | | | | | U | S 19 | 99-P | V156 | 467 | 19990 | 0927 | |
| | | | | | | | | | | U: | s 19 | 99-P | V156 | 236 | 19990 | 0927 | |
| | | | | | | | | | | U: | s 19 | 99-4 | 3643 | 0 | 19993 | 1108 | |
| | | | | | | | | | | U: | s 19 | 99-P | V169 | 100 | 19993 | 1206 | |
| | | | | | | | | | | U | s 19 | 99-P | V173 | 612 | 19993 | 1229 | |
| | | | | | | | | | | U | s 19 | 99-4 | 7443 | 2 | 19993 | 1229 | |
| | | | | | | | | | | U | s 19 | 99-4 | 7638 | 7 | 1999: | 1230 | |
| | | | | | | | | | | U | s · 20 | 00-4 | 9882 | 4 | 20000 | 0204 | |
| | | | | | | | | | | U: | S 20 | 00-5 | 3102 | 5 | 20000 | 0320 | |
| | | | | | | | | | | U | S 20 | 00-P | V197 | 769 | 20000 | 0414 | |
| | | | | | | | | | | U | S 20 | 00-5 | 7822 | 3 | 20000 | 0523 | |

Novel nucleic acid mols. useful as inhibitors of gene expression, AΒ compns., and methods for their use are provided. The invention features novel nucleic acid-based techniques (e.g., enzymic nucleic acid mols. (ribozymes), antisense nucleic acids, 2-5A antisense chimeras, triplex DNA, and antisense nucleic acids contg. RNA-cleaving chem. groups) and their use to modulate the expression of mol. targets impacting the development and progression of cancers, diabetes, obesity, Alzheimer's disease diseases, age-related diseases, and/or hepatitis B infections and related conditions. Catalytic nucleic acids were designed for site-specific cleavage of human mRNA targets encoding protein tyrosine phosphatase 1b, methionine aminopeptidase, .beta.-secretase, presenilin-1, epidermal growth factor receptor-2 (HER2/c-erb2/neu), phospholamban, telomerase, and hepatitis B virus genes. Methods for chem. synthesis of modified nucleoside triphosphates (NTPs) and RNA polymerase-catalyzed incorporation of modified NTPs into catalytic oligonucleotides are also provided. [This abstr. record os one of 6 records for this document necessitated by the large no. of index entries required to fully index the document and publication system constraints.].

L2 ANSWER 9 OF 20 CAPLUS COPYRIGHT 2002 ACS ACCESSION NUMBER: 2001:247142 CAPLUS

DOCUMENT NUMBER:

134:306971

TITLE:

Colon and colon cancer associated cDNAs and proteins and their use in diagnosis and

treatment of colon cancer

INVENTOR(S): Ruben, Steven M.; Barash, Steven C.; Birse,

Charles E.; Rosen, Craig A.

PATENT ASSIGNEE(S): Human Genome Sciences, Inc., USA

SOURCE: PCT Int. Appl., 9787 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE: English FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

APPLICATION NO. PATENT NO. KIND DATE -----_____ WO 2000-US26524 20000928 WO 2001022920 A2 20010405 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG 20010430 AU 2000-77215 20000928 AU 2000077215 A5 P 19990929 US 1999-157137 PRIORITY APPLN. INFO .: P 19991103 US 1999-163280

WO 2000-US26524 W 20000928 AB This invention relates to newly identified colon or colon cancer related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as <scolon cancer antigens<s, and the use of such colon cancer antigens for targeting specific cell types and/or diagnosing, detecting, preventing and treating disorders of the colon, particularly the presence of colon cancer and colon cancer metastases. This invention relates to colon cancer antigens as well as vectors, host cells, antibodies directed to colon cancer antigens and the recombinant or synthetic methods for producing the same. Also provided are diagnostic methods for diagnosing and treating, preventing and/or prognosing disorders related to the colon, including colon cancer, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of colon cancer antigens of the invention. The present invention further relates to inhibiting the prodn. and function of the polypeptides of the present invention.

L2 ANSWER 10 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:832310 CAPLUS

DOCUMENT NUMBER: 135:135386

TITLE: Relationship between apoptosis and expression of

c-met oncogene in gastric carcinomas

AUTHOR(S): Zhuang, Xiaoqiang; Lin, Sanren; Zheng, Jie;

Wang, Lixin; Sun, Guihua; Li, Yan

CORPORATE SOURCE: Department of Gastroenterology, General Hospital

of Guangzhou Command of PLA, Canton, 510010,

Peop. Rep. China

SOURCE: Guangdong Yixue (2000), 21(10), 833-835

CODEN: GUYIEG; ISSN: 1001-9448

PUBLISHER: Guangdongsheng Yixue Qingbao Yanjiuso

DOCUMENT TYPE: Journal LANGUAGE: Chinese

The relationship between expression of c-met oncogene and apoptosis AB in gastric mucosal lesions was studied, and the prognostic significance of gastric carcinomas (GC) was discussed. Expression of c-met was investigated in 145 gastric mucosal lesions by immunohistochem. TUNEL method was used to detect apoptosis. Survival anal. was performed by the long rank test. The expression rates of c-met were 24%, 51%, 62%, 67% and 68%, resp. for chronic superficial gastritis (CSG), chronic atrophic gastritis and intestinal metaplasia (CAC+IM), dysplasia , early GC and advanced GC. The pos. rates were higher in CAG+IM, DYS and GC than that in CSG (P < 0.05). Apoptotic indexes (AI) of the 5 groups were: (4.55 .+-. 2.33)%, (6.43 .+-. 5.60)%, (6.45 .+-.5.12)%, (6.55 .+-. 4.80)%, and %. AI was higher in advanced GC than that in CSG(P < 0.05). Expression of c-met was pos. correlated with AI(P < 0.01). Expression of c-met was also correlated significantly with histol. type, serosal invasion and lymph node metastasis. The expression c-met was significantly higher in Borrmann type 4 GC than that in early GC or in Borrmann type 1,2 (P < 0.01). The survival rate of patients with expression of c-met was significantly lower than that of patients with no expression. The expression of c-met may be assocd. with apoptosis and malignant transformation of gastric mucosa, suggesting that expression of c-met may be a new prognostic factor in gastric carcinoma.

L2 ANSWER 11 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:161492 CAPLUS

DOCUMENT NUMBER: 132:204018

TITLE: Diagnosis and staging of various cancers by

detection of cancer-specific genes (CSG

) and antibody-based treatment

INVENTOR(S): Salceda, Susana; Sun, Yongming; Recipon, Herve;

Cafferkey, Robert

PATENT ASSIGNEE(S): Diadexus Llc, USA

SOURCE: PCT Int. Appl., 58 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2000012758 A1 20000309 WO 1999-US19655 19990901

W: CA, JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,

NL, PT, SE

EP 1109937 A1 20010627 EP 1999-946662 19990901

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI

PRIORITY APPLN. INFO.:

US 1998-98880 P 19980902 WO 1999-US19655 W 19990901

AB The present invention provides a new method for detecting, diagnosing, monitoring, staging, and prognosticating selected cancers including gynecol. cancers such as breast, ovarian, uterine and endometrial cancer and lung cancer by measurement of the levels

of cancer-specific genes (CSG) in cells, tissue, or bodily fluid of a control patient and in a cancer patient, where elevated CSG levels indicated the presence of cancer, and further elevated levels the occurrence of metastasis. Cancer-specific gene sequences are presented which may be used as diagnostic markers for the presence of CSG. Antibodies to these sequences labeled with paramagnetic ions or radioisotopes may be used for imaging the cancer, and antibodies cojugated to cytotoxic agents may be used therapeutically.

REFERENCE COUNT:

REFERENCE(S):

(1) Croce; US 5939258 A 1999 CAPLUS

(2) Paoloni-Giacobno; Genomics 1997, V44, P309

(3) Yu; US 5733748 A 1998 CAPLUS

ANSWER 12 OF 20 CAPLUS COPYRIGHT 2002 ACS L22000:116933 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

132:177721

TITLE:

A novel method of diagnosing, monitoring,

staging, imaging and treating colon

cancer by determining colonspecific genes in body fluids

and tissues

INVENTOR(S):

Sun, Yongming; Recipon, Herve; Macina, Roberto

Α.

PATENT ASSIGNEE(S):

Diadexus Llc, USA PCT Int. Appl., 42 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ______ A1 20000217 WO 1999-US16357 19990720 WO 2000007632

W: CA, JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,

NL, PT, SE

EP 1107798 20010620 EP 1999-937328 A1

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,

PT, IE, FI

PRIORITY APPLN. INFO .:

US 1998-95231 P 19980804 WO 1999-US16357 W 19990720

AB The present invention provides new methods for detecting, diagnosing, monitoring, staging, prognosticating, imaging and treating colon cancer that involves detg. levels of colon-specific gene activity in body fluids and tissues.

REFERENCE COUNT:

REFERENCE(S):

(1) Soppet; US 5861494 1999 CAPLUS

(2) Yu; US 5733748 1998 CAPLUS

ANSWER 13 OF 20 CAPLUS COPYRIGHT 2002 ACS L2

ACCESSION NUMBER:

1999:753379 CAPLUS

DOCUMENT NUMBER:

132:1796

TITLE:

A novel method of diagnosing, monitoring, and

staging colon cancer based

on colon-specific

gene expression

INVENTOR(S):

Macina, Roberto A.; Yang, Fei; Sun, Yongming

PATENT ASSIGNEE(S):

Diadexus Llc, USA PCT Int. Appl., 47 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE PATENT NO. KIND DATE ____ -----_____ WO 9960161 A1 19991125 WO 1999-US10498 19990512

W: CA, JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC,

NL, PT, SE

EP 1080227 20010307 EP 1999-924210 A1

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,

PT, IE, FI PRIORITY APPLN. INFO.:

US 1998-86266 P 19980521 WO 1999-US10498 W 19990512

The present invention provides a new method for detecting, AΒ diagnosing, monitoring, staging, and prognosticating colon cancer vis nine colon-specific

genes (CSGs). Electronic subtractions, transcript imaging and protein functions searches were used to identify clones whose component EST's were exclusively or more frequently found in libraries from specific tumors. Six clones were identified whose expression predominantly occurs in the **colon**, and 1 of these clones was useful as a diagnostic marker for lung cancer.

REFERENCE COUNT:

REFERENCE(S):

(1) Human Genome Sciences Inc; WO 9639419 A1 1996 CAPLUS

ANSWER 14 OF 20 CAPLUS COPYRIGHT 2002 ACS L2

ACCESSION NUMBER: DOCUMENT NUMBER:

1999:495387 CAPLUS 131:154486

TITLE:

Human genes and gene expression products from a colon cancer cell line KM12L4-A cDNA library

INVENTOR(S):

Williams, Lewis T.; Escobedo, Jaime; Innis,

Michael A.; Garcia, Pablo Dominguez;

Sudduth-Klinger, Julie; Reinhard, Christoph; Giese, Klause; Randazzo, Filippo; Kennedy, Giulia C.; Pot, David; Kassam, Altaf; Lamson, George; Drmanac, Radoje; Crkvenjakov, Radomir; Dickson, Mark; Drmanac, Snezana; Labat, Ivan; Leshkowitz, Dena; Kita, David; Garcia, Veronica;

Jones, William Lee; Stache-Crain, Birjit

PATENT ASSIGNEE(S):

Chiron Corporation, USA; Hyseq Inc. PCT Int. Appl., 2479 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.

KIND DATE

DATE APPLICATION NO.

Searcher :

Shears

308-4994

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WO 9938972
                       A2
                            19990805
                                           WO 1999-US1619
                                                            19990128
                            19991223
     WO 9938972
                      A3
            AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
             DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN,
             IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD,
            MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
             SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                            19990816
                                         AU 1999-24716
                                                            19990128
     AU 9924716
                      A1
                                           EP 1999-904288
                                                            19990128
     EP 1053319
                      A2 20001122
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
             PT, IE, FI
                                        US 1998-72910
                                                         Ρ
                                                            19980128
PRIORITY APPLN. INFO.:
                                        US 1998-75954
                                                         Р
                                                            19980224
                                                         Ρ
                                                            19980331
                                        US 1998-80114
                                                         Ρ
                                        US 1998-80515
                                                            19980403
                                                         Р
                                        US 1998-80666
                                                            19980403
                                        US 1998-105234
                                                         P
                                                           19981021
                                                         P 19981027
                                        US 1998-105877
                                                         W 19990128
                                        WO 1999-US1619
AB
     This invention relates to novel human polynucleotides and variants
     thereof, their encoded polypeptides and variants thereof, to genes
     corresponding to these polynucleotides and to proteins expressed by
     the genes. The invention provides the nucleotide sequences for 2502
     human polynucleotides isolated as cDNA clones from the human colon
     cancer cell line KM12L4-A, 2600 validation sequence, plus 146
     sequences assembled as contigs. Many of the cDNA sequences provided
     are differentially expressed in the cancerous state (colon cancer,
     lung cancer, breast cancer) or in specific tissues (e.g., colon).
     Database homol. searches identified various protein families that
     encompass some of the putative protein products. Diagnostic and
     therapeutic agents employing such novel human polynucleotides, their
     corresponding genes or gene products, e.g., these genes and
     proteins, including probes, antisense constructs, and antibodies,
     are also provided.
    ANSWER 15 OF 20 CAPLUS COPYRIGHT 2002 ACS
                         1999:298090 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         131:111015
                         Extract of Solanum muricatum (pepino/CSG
TITLE:
                         ) inhibits tumor growth by inducing apoptosis
                         Ren, Weiping; Tang, Dean G.
AUTHOR(S):
                         Virotech Canada Inc., Windsor, ON, N8W 3K5, Can.
CORPORATE SOURCE:
SOURCE:
                         Anticancer Res. (1999), 19(1A), 403-408
                         CODEN: ANTRD4; ISSN: 0250-7005
                         International Institute of Anticancer Research
PUBLISHER:
DOCUMENT TYPE:
                         Journal
LANGUAGE:
                         English
     Apoptosis, or programmed cell death, is characterized by certain
     distinct morphol. and biochem. features. Most chemotherapeutic
     drugs exert their anti-tumor effects by inducing apoptosis.
     Therefore, an effective compd. inducing apoptosis appears to be a
     relevant strategy to suppress various human tumors. In a search for
     tumor inhibitors from various kinds of plants, we found that exts.
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from Solanum muricatum (CSG) can inhibit tumor growth both in vivo and in vitro by inducing apoptosis. A lyophilized aq. fraction extd. from Solanum muricatum (CSG) was used in The human cell lines tested include: prostate (PC3, this study. DU145), stomach (MKN45), liver (QGY-7721, SK-HEP-1), breast (MDA-MB-435), ovarian (OVCAR), colon (HT29) and lung (NCI-H209) cancer cells; NHP (prostate), HUVEC (umbilical vein endothelial cell), and WI-38 (lung diploid fibroblasts) normal cells. The cell survival was detd. by either Cell Titer MTS cell proliferation kit or trypan blue dye exclusion assay. The apoptosis was analyzed by (a) apoptotic morphol. by light microscopy; (b) DNA ladder formation; (c) PARP cleavage assay. A) CSG possesses selective cytotoxic activity against all the tumor cell lines being tested. The LD50 value is 561-825 .mu.g/mL. B) CSG showed a much lower cytotoxicity to NHP, HUVEC and WI-38 normal cell lines with LD50 value being 2.8-3.2 mg/mL, which is 3-6 fold higher than on tumor cells. C) The in vivo study demonstrated that injection of CSG (100 .mu.g) directly into tumor mass can reduce the tumor vol. dramatically in nude mice inoculated with MKN45 gastric cancer cells. D) CSG-mediated tumor growth inhibition is through induction of apoptotic cell death, as manifested by (a) typical apoptotic morphol.; (b) DNA ladder formation; and (c) PARP cleavage assay. Taken together, the present study suggests, for the first time, that CSG may represent promising new chem. entity which preferentially targets various tumor cells by triggering apoptosis.

REFERENCE COUNT:

REFERENCE(S):

17

- (2) Chiang, H; Anticancer Res 1991, V11, P1911 CAPLUS
- (4) Hickman, J; Cancer Metastasis Rev 1992, V11, P121 CAPLUS
- (5) Hsu, S; Biochem Biophys Res Com 1996, V229, P1 CAPLUS
- (7) Mohanan, P; Cancer Lett 1996, V110, P71 CAPLUS
- (8) Mohanan, P; Cancer Lett 1997, V112, P219 CAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 16 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1998:794684 CAPLUS

DOCUMENT NUMBER:

130:221220

TITLE:

Significance of CD44S and CD44V6 expression in

gastric carcinoma

AUTHOR(S):

Li, Xueyan; Hu, Jialu

CORPORATE SOURCE:

Department of Digestive Medicine, 4th Military

Medical University Xijing Hospital, Xi'an,

710033, Peop. Rep. China

SOURCE:

Disi Junyi Daxue Xuebao (1998), 19(5), 534

CODEN: DJDXEG; ISSN: 1000-2790

PUBLISHER:

Disi Junyi Daxue Xuebao Bianjibu

DOCUMENT TYPE:

Journal

LANGUAGE:

Chinese

AB CD44 is a surface adhesion protein, its variants were assocd. with tumor metastasis. CD44S and CD44V6 expression were examd. by immunohistochem. staining in 54 benign gastric disease including 33 chronic superficial gastritis (CSG), 21 chronic atrophic gastritis (CAG) accompanied intestinal metastasis

(IM), and 63 gastric carcinoma (GC) including 35 cases with, and 28 cases without lymph node metastasis. The pos. expression of CD44S in CSG, CAG/IM, and GC were 45.5, 57.1, and 52.4%; and CD44V6 were 0, 19.1, and 53.9% resp. CD44S expression in intestinal type gastric carcinoma and diffused type gastric carcinoma were 55.3 and 48%, P> 0.05; while CD44V6 expression were 73.7 and 24%, P< 0.05. CD44S and CD44V6 expression were not related with gastric carcinoma histol. type and tumor size. CD44S expression in gastric carcinoma with or without metastasis were 46.7 and 60.7%, P> 0.05; while CD44V6 expression were 68.8 and 35.7%, P< 0.05. The CD44V6 expression was significantly higher in intestinal type and lymph node metastasized gastric carcinoma. The results suggest that the CD44S and CD44V6 expression might be indexes in evaluation and prediction of lymph node metastasis of gastric carcinoma.

ANSWER 17 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1998:202636 CAPLUS

DOCUMENT NUMBER:

128:240996

TITLE:

Human colon-specific cDNA and protein

sequences and use as diagnostic markers for

colon cancer presence and

metastasis

INVENTOR(S):

Yu, Guo-Liang; Rosen, Craig

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., USA

SOURCE:

U.S., 50 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. [| DATE |
|----------------------|------|----------|---------------------|----------|
| | | | | |
| US 5733748 | Α | 19980331 | US 1995-469667 1 | L9950606 |
| US 6337195 | В1 | 20020108 | US 1998-224110 1 | L9980331 |
| PRIORITY APPLN. INFO | .: | | US 1995-469667 A3 1 | 19950606 |

Human colon specific gene polypeptides AΒ and DNA (RNA) encoding such polypeptides are claimed, along with procedures for producing these polypeptides by recombinant techniques, their use as diagnostic markers for colon cancer presence and progression, antibodies to the polypeptides which may be used as a vaccine, and methods for screening for agonists and antagonists which may have therapeutic use.

ANSWER 18 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1997:105242 CAPLUS

DOCUMENT NUMBER:

126:114205

TITLE:

Human colon-specific

genes and proteins

INVENTOR(S):

Yu, Guo-Liang; Rosen, Craig A.

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., USA; Yu, Guo-Liang;

Rosen, Craig A.

SOURCE:

PCT Int. Appl., 79 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

Searcher : 308-4994 Shears

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

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KIND DATE
                                         APPLICATION NO.
                                                           DATE
    PATENT NO.
                           -----
                                         ______
                     ____
                                    WO 1995-US7289 19950606
    WO 9639419
                           19961212
                    A1
        W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI,
            GB, GE, HU, JP, KE, KG, KP, KR, KZ, LK, LT, LU, LV, MD, MG,
            MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT,
            UA, US, UZ, VN
        RW: KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE,
            IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML,
            MR, NE, SN, TD, TG
                           19961212
                                          CA 1995-2221798 19950606
    CA 2221798
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                           19961224
                                         AU 1995-28205
                                                           19950606
    AU 9528205
                      A1
                           19980617
                                         EP 1995-923764
                                                          19950606
    EP 847398
                     A1
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,
            PT, IE
                           19990608
                                          JP 1995-500380
                                                          19950606
    JP 11506342
                      T2
                                       WO 1995-US7289
                                                          19950606
PRIORITY APPLN. INFO.:
    Thirteen human colon-specific cDNAs and their deduced amino acid
    sequences and procedures for producing such polypeptides by
    recombinant techniques are provided. Two of the cDNAs are
    full-length. Also disclosed are methods for utilizing such
    polypeptides or polypeptides as a diagnostic marker for
    colon cancer and as an agent to det. if
    colon cancer has metastasized. Also
    disclosed are antibodies specific to the colon-
    specific gene polypeptides which may be used to
    target cancer cells and be used as part of a colon
    cancer vaccine. Methods of screening for agonists and
    antagonists for the polypeptides and therapeutic uses of the
    antagonists are disclosed.
```

L2 ANSWER 19 OF 20 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER:

1995:200493 CAPLUS

DOCUMENT NUMBER:

122:7233

TITLE:

A gene expressed in colon mucosa gene that is expressed at lower levels in colon adenomas and

adenocarcinomas

INVENTOR(S):

Schweinfest, Clifford W.; Papas, Takis S. United States Dept. of Health and Human

Services, USA

SOURCE:

PCT Int. Appl., 44 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT ASSIGNEE(S):

| PAT | ENT NO. | | KINI | D DATE | | APPL] | CATION NO. | DATE | |
|-----|---------|-----|-------|-------------|-----|---------|------------|------------|-----|
| | | | | | | | | | |
| WO | 9420616 | | A1 | 19940915 | | WO 19 | 994-US1860 | 19940304 | |
| | W: AU, | CA, | JP | | | | | | |
| | RW: AT, | BE, | CH, I | DE, DK, ES, | FR, | GB, GR, | IE, IT, L | J, MC, NL, | PT, |
| | SE | | | | | | | | |
| AU | 9463508 | | A1 | 19940926 | | AU 19 | 994-63508 | 19940304 | |
| US | 5569755 | | Α | 19961029 | | US 19 | 995-424567 | 19950417 | |

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US 5831015
                      Α
                           19981103
                                          US 1996-711928
                                                           19960911
    US 6210887
                      В1
                           20010403
                                          US 1998-184937
                                                          19981102
PRIORITY APPLN. INFO.:
                                       US 1993-26045 A 19930305
                                       WO 1994-US1860
                                                      W 19940304
                                                       A3 19950417
                                       US 1995-424567
                                       US 1996-711928
                                                      A3 19960911
    A new gene called DRA, for down regulated in adenoma, is expressed
AΒ
    at lower levels in colon adenomas than in normal tissues,
    maps to chromosome 7 and is believed to encode a tumor
    suppressor. The DRA gene encodes a highly hydrophobic protein with
    charged clusters located primarily in the carboxyl terminus. The
    mRNA appears to be strictly limited to the mucosa of normal colon
    and it is down-regulated early in colon tumorigenesis. Absence of
    the DRA polypeptide in tissue that usually expresses it can be used
    as an indicator of tissue abnormality. The DRA gene and cDNA may
    also have therapeutic uses. A cDNA from the gene was cloned by
    differential screening of banks from normal colon and colon
    adenocarcinoma.
    ANSWER 20 OF 20 CAPLUS COPYRIGHT 2002 ACS
L2
ACCESSION NUMBER:
                        1993:78665 CAPLUS
                        118:78665
DOCUMENT NUMBER:
                        Inherited and somatic mutations of the APC gene
TITLE:
                        associated with colorectal cancer of humans
                        Kinzler, Kenneth W.; Vogelstein, Bert; Anand,
INVENTOR(S):
                        Rakesh; Hedge, Philip John; Markham, Alexander
                        Fred; Albertsen, Hans; Carlson, Mary L.; Groden,
                        Joanna L.; Joslyn, Geoff; et al.
                        Johns Hopkins University, USA; Imperial Chemical
PATENT ASSIGNEE(S):
                        Industries PLC; University of Utah; Cancer
                        Institute
SOURCE:
                        PCT Int. Appl., 138 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                                           DATE
    PATENT NO.
                     KIND DATE
                                         APPLICATION NO.
                          _____
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                                        WO 1992-US376 19920116
                    A1 19920806
    WO 9213103
        W: AT, AU, BB, BG, BR, CA, CH, CS, DE, DK, ES, FI, GB, HU, JP,
            KP, KR, LK, LU, MG, MN, MW, NL, NO, PL, RO, RU, SD, SE
        RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FR, GA, GB,
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GN, GR, IT, LU, MC, ML, MR, NL, SE, SN, TD, TG
                                        US 1991-741940
                                                         19910808
                          19941004
    US 5352775
                     Α
                          19920827
                                         AU 1992-13669
                                                         19920116
    AU 9213669
                     A1
                                         EP 1992-906080
                                                         19920116
    EP 569527
                     A1
                          19931118
                    В1
                         20010314
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, MC, NL, SE
                                                        19920116
    JP 07500241
                    T2 19950112
                                         JP 1992-506203
                                         AT 1992-906080
                                                         19920116
    AT 199746
                      E
                          20010315
                                                     A 19910116
PRIORITY APPLN. INFO.:
                                      GB 1991-963
                                                     A 19910808
                                      US 1991-741940
                                                      A 19910116
                                      GB 1991-962
                                                      A 19910116
                                      GB 1991-974
                                                      A 19910116
                                      GB 1991-975
                                                      A 19920116
                                      WO 1992-US376
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A human gene that shows inherited and somatic mutations assocd. with AB colorectal cancer is cloned and characterized. The gene and its product are useful as markers in the diagnosis and prognosis of the disease. A series of YAC clones of the 5q21 region were cloned by screening with markers for the region. Six genes expressed in normal colon cells and in colorectal, lung and bladder tumors were found in the region. These genes were: the FER gene at 5q11-23 similar to the v-abl gene; TB1 showing some similarity to brown adipose tissue uncoupling proteins; MCC and TB2; and APC. A cDNA from the APC gene had an open reading frame of 8,535 nucleotides that encoded a protein with some similarity to myosins and intermediate filament proteins and to to the ral2 gene product of yeast. The assocn. of these genes and mutant alleles with colorectal cancer was studied by std. methods. The gene that showed the greatest no. of germline and somatic mutations was APC and the characterization of a no. of the mutations is described.

(FILE OMEDIINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JACST-EPLOS, JAPIO, CANCERLIT' ENTERED AT 10:08:11 ON 18 JAN 2002) 28 S L2

24 DOP REM L3 (4 DOPLICATES REMOVED)

DERWENT INFORMATION LTD ANSWER 1 OF 24 WPIDS COPYRIGHT 2002

ACCESSION NUMBER:

2001-616504 [71] WPIDS

DOC. NO. NON-CPI:

N2001-459822

DOC. NO. CPI:

C2001-184647

TITLE:

New colon cancer specific

polypeptides and polynucleotides, useful for

detecting, diagnosing, monitoring, staging, imaging

and treating cancers, particularly

colon cancer.

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

HU, P; MACINA, R A; PIDERIT, A; RECIPON, H; YANG, F

(DIAD-N) DIADEXUS INC

COUNTRY COUNT:

PATENT ASSIGNEE(S): PATENT INFORMATION:

> PATENT NO KIND DATE WEEK LA PG ------

23

WO 2001073030 A2 20011004 (200171) * EN 105

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR W: AU CA JP US

APPLICATION DETAILS:

APPLICATION DATE PATENT NO KIND _____ WO 2001-US9737 20010326 WO 2001073030 A2

PRIORITY APPLN. INFO: US 2000-192667P 20000328

2001-616504 [71] WPIDS

AB WO 200173030 A UPAB: 20011203

NOVELTY - An isolated colon cancer specific gene

(CSG) polynucleotide (I) comprising:

(a) one of 57 sequences (S1) of defined base pairs (bp) as given in specification;

(b) its fragment of 15 contiguous nucleobases;

Shears 308-4994 Searcher :

```
(c) a nucleic acid sequence which, due to degeneracy in genetic
coding, has variations in (S1); or
     (d) a nucleic acid sequence which hybridizes under stringent
conditions to an antisense sequence of (S1), is new.
     DETAILED DESCRIPTION - An isolated colon
cancer specific gene (CSG) polynucleotide (I)
comprising:
     (a) one of 57 sequences (S1) of defined base pairs (bp) as
given in specification such as 523, 528, 478, 495, 455, 489, 545,
220, 484, 350, 322, 306, 143, 508, 582, 582, 521, 244 and 600 bp;
     (b) its fragment of 15 contiguous nucleobases;
     (c) a nucleic acid sequence which, due to degeneracy in genetic
coding, has variations in (S1); or
     (d) a nucleic acid sequence which hybridizes under stringent
conditions to an antisense sequence of (S1), is new.
     INDEPENDENT CLAIMS are also included for the following:
     (1) an antisense oligonucleotide (II) which hybridizes to (I);
     (2) a vector (III) comprising (I);
     (3) a host cell (IV) comprising (III);
     (4) a CSG polypeptide (V) encoded by (I);
     (5) producing (V);
     (6) producing a cell expressing (V) by transforming or
transfecting a cell with (III) so that the cell under appropriate
culture conditions, expresses (V);
     (7) an antibody (VI) which is immunospecific for (V);
     (8) a colon cancer specific gene (
CSG) for diagnosing colon cancer,
comprising (I) or (V);
     (9) a CSG polypeptide agonist or antagonist
identified using (V); and
     (10) a vaccine (VII) comprising (V) or a vector expressing (V)
which induces an immune response against (V) in a mammal.
     ACTIVITY - Cytostatic.
    MECHANISM OF ACTION - Vaccine; gene therapy. No supporting data
     USE - CSG is useful for diagnosing, staging,
monitoring colon cancer for onset of
metastasis or a change in stage of colon
cancer, diagnosing metastases of colon
cancer in a patient, by determining levels of CSG
in a sample of cells, tissues, or body fluids and comparing it with
levels of CSG in normal human control, where an increase
in determined CSG level is associated with cancer
. CSG is also useful for identifying potential therapeutic
agents for use in imaging and treating colon
cancer, by screening molecules for ability to bind to
CSG. (V) is useful for identifying compounds which
antagonize or agonize the CSG polypeptide, by contacting
cells or cell membrane which express (V) with a candidate compound
and monitoring the cells for changes in CSG polypeptide
activities or binding as compared to cells or cell membranes not
contacted with the candidate compound. (VI) labeled with
paramagnetic ions or a radioisotope is useful for imaging
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claimed). (I), (V) and (VI) are useful for detecting the effect of

colon cancer and (VI) conjugated to a cytotoxic
agent is useful for treating colon cancer. (VII)
is useful for inducing an immune response against CSG

polypeptide and treating colon cancer (all

added compounds on the production of **CSG** mRNA and polypeptides in cells. (V) is also useful to identify membrane bound or soluble receptors. (VI) is useful to isolate or identify clones expressing **CSG** polypeptide and to purify the polypeptides by affinity chromatography. Dwg.0/0

L4 ANSWER 2 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2001-389934 [41] WPIDS

DOC. NO. CPI: C2001-118811

TITLE: Novel cancer specific gene and its protein useful

for detecting, diagnosing, monitoring, staging, prognosticating, imaging and treating prostate

cancer.

DERWENT CLASS: B04 D16

INVENTOR(S): ALI, S; CAFFERKEY, R; RECIPON, H; SUN, Y

PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS INC

COUNTRY COUNT: 21

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2001039798 A1 20010607 (200141)* EN 52

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE TR

W: CA JP

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|--------------|-------|----------------|------------|
| | | | |
| WO 200103979 | 98 A1 | WO 2000-US3292 | 7 20001205 |

PRIORITY APPLN. INFO: US 1999-169083P 19991206

AN 2001-389934 [41] WPIDS

AB WO 200139798 A UPAB: 20010724

NOVELTY - A cancer specific gene (CSG) (I) comprising a sequence (S1) of 310, 2994, 230, 660, 191 or 647 nucleotides (nts) fully defined in the specification or its variant, a protein or its variant expressed by S1, or a polynucleotide which is capable of hybridizing under stringent conditions to an antisense sequence of S1, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) diagnosing (M1) the presence of prostate cancer in a patient, by determining levels of CSG in samples (Sa1) such as cells, tissues or bodily fluids in a patient, and comparing the determined levels of CSG with levels of CSG in samples (Sa2) such as cells, tissues or bodily fluids from a normal human control;
- (2) diagnosing (M2) metastases of prostate cancer in a patient, by identifying a patient having prostate cancer that is not known to have metastasized, determining CSG levels in Sal, and comparing the determined CSG levels with levels of CSG in Sa2, where an increase in determined CSG levels in the patient versus normal human control is associated with a cancer which has metastasized;
 - (3) staging (M3) prostate cancer in a patient, by identifying a

patient having prostate cancer, determining CSG levels in Sa1, and comparing the determined CSG levels with levels of CSG in Sa2, where an increase in determined CSG levels in the patient versus the normal human control is associated with a cancer which is progressing and a decrease in the determined CSG levels is associated with a cancer which is regressing or in remission;

- (4) monitoring (M4) prostate cancer in a patient for the onset of metastasis, by identifying a patient having prostate cancer that is not known to have metastasized, periodically determining CSG levels in Sal, and comparing them with levels of CSG in Sa2, where an increase in any one of the determined CSG levels in the patient versus normal human control is associated with a cancer which has metastasized;
- (5) monitoring (M5) a change in stage of prostate cancer in a patient, by identifying a patient having prostate cancer, periodically determining CSG levels in Sal fluids from the patient, and comparing them with levels of CSG in Sa2, where an increase in any one of the determined CSG levels in the patient versus normal human control is associated with a cancer which is progressing in stage and a decrease is associated with a cancer which is regressing in stage or in remission;
- (6) identifying (M6) potential therapeutic agents for use in imaging and treating prostate cancer, by screening molecules for an ability to bind to CSG, which is indicative of the molecule being useful in imaging and treating prostate cancer;
 - (7) an antibody (Ab) which specifically binds (I);
- (8) treating (M7) prostate cancer in a patient, by administering (Ab), or a molecule which downregulates expression or activity of a CSG;
 - (9) imaging (M8) prostate cancer by administering (Ab); and
 - (10) a vaccine for treating prostate cancer comprising (I). ACTIVITY - Cytostatic.

MECHANISM OF ACTION - Gene therapy; vaccine. No supporting data given.

USE - Ab is useful for imaging and treating prostate cancer in a patient. CSG protein is useful for inducing an immune response against a target cell expressing a CSG (claimed).

(I) is useful as diagnostic marker for detecting, diagnosing (metastases and disease), monitoring (cancer and changes in cancer), staging, prognosticating, imaging and treating prostate cancer (all claimed).

ADVANTAGE - The method is more sensitive and accurate for staging human cancer. $\ensuremath{\mathsf{Dwg.0/0}}$

L4 ANSWER 3 OF 24 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER:

2001189505 EMBASE

TITLE:

hTERT expression and cellular immunity in gastric

cancer and precancerosis.

AUTHOR:

Yao X.X.; Yin L.; Zhang J.Y.; Bai W.Y.; Li Y.M.; Sun

Z.C.

CORPORATE SOURCE:

Dr. X.X. Yao, Department of Digestive Medicine, 2nd Hosp. of Hebei Med. University, Shijiazhuang 050000,

Hebei Province, China. Yaoxixian@263.net

SOURCE:

World Chinese Journal of Digestology, (2001) 9/5

(508-512).

Refs: 51

ISSN: 1009-3079 CODEN: SHXZF2

COUNTRY:

China

DOCUMENT TYPE: FILE SEGMENT:

Journal; Article

016 Cancer 026

Immunology, Serology and Transplantation

048 Gastroenterology

LANGUAGE:

Chinese

SUMMARY LANGUAGE:

English; Chinese

AIM To observe the expression of human telomerase reverse transcriptase (hTERT) in gastric carcinomas and precancerous lesions, evaluate the immune state of such patients, and to study the clinical implications of hTERT and immune state for the diagnosis, treatment and prognosis of gastric cancer. METHODS In situ hybridization was used to detect the expression of hTERT mRNA in 116 endoscopic biopsies of gastric mucosa. Tissue samples were analyzed as follows: 30 cases of chronic superficial gastritis (CSG), 44 of precancerous lesions (including 27 of chronic atrophy gastritis, 8 of hyperplastic ploy and 9 of gastric ulcer) and 42 of gastric cancer (GC). At mean time, the T lymphocyte subsets (CD3(+), CD4(+), CD8(+), CD4(+) / CD8(+)) and natural killer cell (NK) in peripheral blood were determined by flow cytometrical analysis (FCM) in 30 cases of CSG, 27 of precancerosis (chronic atrophy gastritis, CAG), 42 of GC and the data were campared with those of normal controls (NC). RESULTS The positive rate of hTERT varied as follows: 0%(0/30) in CSG, 36% (16/44) in precancerous lesions and 86% (36/42) in GC. The expression of hTERT mRNA was not associated with patients gender, tumor location, macroscopic type, lymph node metastasis and degree of differentiation. The CD3(+) and CD4(+) of CSG were lower than that of NC (P<0.05). Meanwhile, the T lymphocyte subsets (CD3(+), CD4(+), CD4(+) / CD8(+) ratio) were remarkably lower than that of NC and CSG (P<0.05-0.01). Values of T cells and NK cells of GC group were abnormal significantly as compared with CAG (P<0.05-0.01). Furthermore, with the tumor progression, the function of T cells was weakened gradually. CONCLUSION The expression of telomerase may be a crucial step in gastric carcinogenesis and increased hTERT mRNA may serve as a novel marker for diagnosis of gastric cancer. The immune state of patients with gastric cancer and precancerosis was somewhat depressed, which indicates the importance of cellular immunity in cancer patients.

ANSWER 4 OF 24 BIOSIS COPYRIGHT 2002 BIOSIS

ACCESSION NUMBER: DOCUMENT NUMBER:

2001:114238 BIOSIS PREV200100114238

TITLE:

Colon specific gene and

protein.

AUTHOR(S):

Soppet, Daniel R.; Li, Yi; Dillon, Patrick J. (1)

(1) Gaithersburg, MD USA CORPORATE SOURCE:

ASSIGNEE: Human Genome Sciences, Inc.

PATENT INFORMATION: US 6080722 June 27, 2000

SOURCE:

Official Gazette of the United States Patent and Trademark Office Patents, (June 27, 2000) Vol. 1235,

No. 4, pp. No Pagination. e-file.

ISSN: 0098-1133.

DOCUMENT TYPE:

Patent English

LANGUAGE:

Human colon specific gene polypeptides

308-4994 Searcher : Shears

and DNA (RNA) encoding such polypeptides and a procedure for producing such polypeptides by recombinant techniques is disclosed. Also disclosed are methods for utilizing such polynucleotides or polypeptides as a diagnostic marker for colon cancer and as an agent to determine if colon cancer has metastasized. Also disclosed are antibodies specific to the colon specific gene polypeptides which may be used to target cancer cells and be used as part of a colon cancer vaccine. Methods of screening for agonists and antagonists for the polypeptide and therapeutic uses of the antagonists are also disclosed.

L4 ANSWER 5 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-339531 [29] WPIDS

DOC. NO. NON-CPI: N2000-254921 DOC. NO. CPI: C2000-103001

TITLE: Diagnosing, staging and monitoring the presence and

metastases of prostate cancer especially useful for treating prostate cancer comprises measuring changes in cancer specific gene levels.

DERWENT CLASS: B04 D16 S03

INVENTOR(S): CAFFERKEY, R; RECIPON, H; SALCEDA, S

PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS INC; (DIAD-N) DIADEXUS LLC

COUNTRY COUNT: 22

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000023111 A1 20000427 (200029)* EN 74

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1131095 A1 20010912 (200155) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO K | IND | | PLICATION | DATE |
|-----------------------------|-----|----------|---|----------------------|
| WO 2000023111 EP 1131095 | | WO EP | 1999-US24331 1999-955004 1999-US24331 | 19991019 19991019 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|--------------|
| | | |
| EP 1131095 | Al Based on | WO 200023111 |

PRIORITY APPLN. INFO: US 1998-104737P 19981019

AN 2000-339531 [29] WPIDS

AB WO 200023111 A UPAB: 20000617

NOVELTY - A method for diagnosing the presence of prostate cancer in a patient, comprising determining levels of cancer specific genes (CSG) in cells, tissues or bodily fluids, and comparing the determined levels of CSG with levels of CSG from a normal human control, is new. A change in determined levels of CSG in the patient versus the control is associated with the

presence of prostate cancer.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a method of diagnosing **metastases** of prostate cancer in a patient comprising:
- (a) identifying a patient having prostate cancer that is not known to have metastasized;
- (b) determining CSG levels in a sample of cells, tissues, or bodily fluid from the patient; and
- (c) comparing the determined CSG levels with CSG levels of a normal human control, where an increase in CSG levels in the patient versus the control, is associated with a cancer which has metastasized;
- (2) a method of staging prostate cancer in a patient having prostate cancer, comprising:
 - (a) identifying a patient having prostate cancer;
- (b) determining CSG levels in a sample of cells, tissue, or bodily fluid from the patient; and
- (c) comparing determined CSG levels with CSG levels of a normal human control, where an increase in CSG levels in the patient versus the control is associated with a progressing cancer, and a decrease in the CSG levels is associated with a regressing cancer;
- (3) a method of monitoring prostate cancer in a patient for the onset of **metastasis** comprising:
- (a) identifying a patient having prostate cancer that is not known to have metastasized;
- (b) periodically determining CSG levels in samples of cells, tissues, or bodily fluid from the patient; and
- (c) comparing the CSG levels with CSG levels of CSG of a normal human control, where an increase in any one of the periodically determined CSG levels in the patient versus the control is associated with a cancer which has metastasized;
- (4) a method of monitoring a change in stage of prostate cancer in a patient comprising:
 - (a) identifying a patient having prostate cancer;
 - (b) periodically determining levels of CSG;
- (c) comparing the CSG levels with CSG levels of a normal human control, where an increase in any one of the periodically determined CSG levels in the patient versus the control is associated with a progressing cancer, and a decrease is associated with a regressing cancer;
- (5) a method of identifying potential therapeutic agents for use in imaging and treating prostate cancer, comprising screening molecules for an ability to bind to CSG, which indicates the molecule is useful in imaging and treating prostate cancer;
 - (6) an antibody which specifically binds CSG; and
- (7) a method of imaging or treating prostate cancer in a patient by administering an antibody of (7).

ACTIVITY - Cytostatic.

MECHANISM OF ACTION - The antibody, conjugated to a cytotoxic agent, binds to cancer specific genes, in vivo.

USE - The method is useful for diagnosing, staging and monitoring the presence and **metastases** of cancer (claimed). The antibodies which specifically binds **CSG** or fragments of such antibodies can be used in treating prostate cancer, and to detect or image, localization of **CSG** in a

patient in order to diagnose a disease or condition (claimed). The antibodies may also be used in the treatment of diseases characterized by the expression of CSG.

ADVANTAGE - The new method provides an earlier diagnosis for the presence and **metastasis** of prostate cancer, which significantly increase the chances of a cure. It provides a sensitive method for diagnosing, and staging, prostate cancer to determine if the cancer has **metastasized**, and for monitoring the progress or stage of the disease, which has not **metastasized**.

Dwg.0/0

L4 ANSWER 6 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

2000-339528 [29] WPIDS

DOC. NO. NON-CPI:

N2000-254918

DOC. NO. CPI:

C2000-102998

TITLE:

Diagnosing, detecting, staging, monitoring, imaging

and treating cancers, especially useful for detecting prostate cancer comprises measuring changes in levels of cancer specific genes in

cells, tissues and body fluids.

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

ALI, S M; CAFFERKEY, R; RECIPON, H; SALCEDA, S;

SUN, Y

PATENT ASSIGNEE(S):

(DIAD-N) DIADEXUS INC; (DIAD-N) DIADEXUS LLC

COUNTRY COUNT:

22

PATENT INFORMATION:

| PATENT NO | O KIND | DATE | WEEK | LA | PG |
|-----------|--------|------|------|----|----|
| | | | | | |

WO 2000023108 A1 20000427 (200029)* EN 33

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1126877 A1 20010829 (200150) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO K | IND | | PLICATION | DATE |
|---------------|-----|----|--------------|----------|
| | | | | |
| WO 2000023108 | A1 | WO | 1999-US23764 | 19991018 |
| EP 1126877 | A1 | EΡ | 1999-954867 | 19991018 |
| | | WO | 1999-US23764 | 19991018 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|--------------|
| | | |
| EP 1126877 | Al Based on | WO 200023108 |

PRIORITY APPLN. INFO: US 1998-104741P 19981019

AN 2000-339528 [29] WPIDS

AB WO 200023108 A UPAB: 20000617

NOVELTY - A method for diagnosing the presence of prostate cancer, comprising measuring levels of cancer specific genes (CSG) in cells, tissues or bodily fluids, and comparing the measured CSG levels with levels from a normal human control, where a change in measured CSG levels in the patient versus the

control is associated with the presence of prostate cancer.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a method for diagnosing the **metastases** of prostate cancer in a patient comprising:
- (a) identifying a patient having prostate cancer that is not known to have **metastasized**;
- (b) measuring CSG levels in cells, tissues or bodily fluid of the patient; and
- (c) comparing the measured CSG levels with levels of a normal human control, where an increase in measured CSG levels in the patient versus the control is associated with a cancer which has metastasized;
- (2) a method for staging prostate cancer in a patient having prostate cancer comprising:
 - (a) identifying a patient having prostate cancer;
- (b) measuring CSG levels in cells, tissues or bodily fluid of the patient; and
- (c) comparing the measured **CSG** levels with levels of a normal human control, where an increase in measured **CSG** levels in the patient versus the control is associated with a cancer which is progressing, and a decrease in the measured **CSG** levels is associated with a cancer which is regressing or in remission;
- (3) a method of monitoring prostate cancer in a patient for the onset of metastasis comprising:
- (a) identifying a patient having prostate cancer that is known to have metastasized;
- (b) periodically measuring levels of CSG in samples of cells, tissues or bodily fluid from the patient;
- (c) comparing the CSG levels with levels of a normal human control, where an increase in any one of the periodically measured CSG levels in the patient versus the control is associated with a cancer which has metastasized;
- (4) a method of monitoring a change in stage of prostate cancer in a patient comprising:
 - (a) identifying a patient having prostate cancer;
- (b) periodically measuring levels of CSG in samples of cells, tissues or bodily fluid from the patient;
- (c) comparing the CSG levels with levels of a normal human control, where an increase in any one of the periodically measured CSG levels in the patient versus the control is associated with a progressing cancer, and a decrease is associated with a regressing cancer.
 - (5) an antibody which specifically binds CSG; and
- (6) a method of imaging or treating prostate cancer in a patient, comprising administering the antibody of (6).

ACTIVITY - Cytostatic.

MECHANISM OF ACTION - The antibody, conjugated to a cytotoxic agent, binds to cancer specific genes.

USE - The method is useful for diagnosing, detecting, staging, monitoring, and imaging for the presence and **metastases** of prostate cancer. The antibodies which specifically bind to CSG may be used to detect or image localization of CSG in a patient in order to detect or diagnose a disease or condition, and to treat prostate cancer. All claimed.

ADVANTAGE - The new method provides an earlier diagnosis for the presence and metastasis of prostate cancer, which

significantly increase the chances of cure. It provides a sensitive method for diagnosing and staging of prostate cancer to determine whether or not such cancer has **metastasized**, and for monitoring the progress of the disease, which has not **metastasized** for the onset of **metastasis**.

Dwg.0/0

L4 ANSWER 7 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-328946 [28] WPIDS

DOC. NO. NON-CPI: N2000-247638 DOC. NO. CPI: C2000-099678

TITLE: Detecting, diagnosing and monitoring

gastrointestinal cancers comprises measuring the levels of cancer specific gene/protein 2 (CC2) in

tissues or bodily fluids.

DERWENT CLASS: B04 D16 S03 INVENTOR(S): MACINA, R A

PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS LLC

COUNTRY COUNT: 22

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000020640 A1 20000413 (200028)* EN 33

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1117833 A1 20010725 (200143) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO KIN | D API | PLICATION | DATE |
|-----------------|-------|-----------------------------|----------------------|
| WO 2000020640 A | | 1999-US22725 | |
| EP 1117833 A | | 1999-950047 1999-US22725 | 19990930 19990930 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|--------------|
| | | |
| EP 1117833 | A1 Based on | WO 200020640 |

PRIORITY APPLN. INFO: US 1998-102879P 19981002

AN 2000-328946 [28] WPIDS

AB WO 200020640 A UPAB: 20000613

NOVELTY - Diagnosing the presence of gastrointestinal cancer (GC), comprising measuring a change in levels of cancer specific gene/protein 2 (CC2) in cells, tissues or bodily fluids in a patient compared with CC2 levels in a normal human control, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) diagnosing metastases of a GC in a patient, comprising:
- (a) identifying a patient having a GC that is not known to have metastasized; and
- (b) the above new method. where an increase in measured CC2 levels in the patient is associated with a cancer which has

metastasized;

- (2) staging a GC in a patient having a GC, comprising steps (a)-(b) of method of (1), where an increase in CC2 levels in the patient is associated with a cancer which is progressing and a decrease is associated with a cancer which is regressing or in remission;
- (3) monitoring a change in the stage of a GC in a patient, comprising step (a) of the method of (1) and:
- (a) periodically measuring the level of CC2 in samples of cells, tissues or bodily fluids from the patient; and
- (b) as for step (c) of the method of (1), wherein an increase in CC2 levels in the patient is associated with a cancer which has metastasized/is progressing and a decrease is associated with a cancer which is regressing or in remission;
 - (4) an antibody that specifically binds CC2;
- (5) imaging a GC cancer in a patient, comprising administering the antibody of (4) (which is preferably labeled with paramagnetic ions or a radioisotope) to the patient; and
- (6) a method of treating a GC in a patient, comprising administering the antibody of (5) (which is preferably conjugated to a cytotoxic agent) to the patient.

USE - The methods are used for diagnosing the presence of gastrointestinal cancers such as stomach cancer, cancer of the small intestine, and colon cancer, especially for a gastrointestinal cancer which has not metastasized. The methods may also be used for staging and monitoring gastrointestinal cancer. Antibodies which specifically bind to colon specific gene 2 (CC2) can also be used in vivo in patients suspected of having gastrointestinal cancers, for treatment and imaging (all claimed).

ADVANTAGE - The new methods are sensitive and specific and allow for early diagnosis of gastrointestinal cancer. This means that treatment can commence earlier. Furthermore, the methods are not invasive, unlike prior art surgical procedures. Dwg.0/0

L4 ANSWER 8 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-283453 [24]

DOC. NO. NON-CPI: N2000-213335 DOC. NO. CPI: C2000-085572

TITLE: Methods for diagnosing, staging, imaging and

treating gynecologic and testicular cancers by measuring expression of a cancer specific gene.

WPIDS

DERWENT CLASS: B04 D16 S03

INVENTOR(S): ALI, S M; CAFFERKEY, R
PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS LLC

COUNTRY COUNT: 22

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000016805 A1 20000330 (200024)* EN 33

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1115426 A1 20010718 (200142) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO KI | IND | API | PLICATION | DATE |
|-----------------------------|----------|-----|---|----------|
| WO 2000016805 EP 1115426 | A1 A1 | EP | 1999-US21774 1999-948349 1999-US21774 | 19990923 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|--------------|
| | | |
| EP 1115426 | Al Based on | WO 200016805 |

PRIORITY APPLN. INFO: US 1998-101522P 19980923

AN 2000-283453 [24] WPIDS

AB WO 200016805 A UPAB: 20000522

NOVELTY - Methods ((I) - (IV)) for diagnosing, staging, imaging and treating gynecologic and testicular cancers by measuring expression of a cancer specific gene (CSG) (comprising a defined 1081 nucleotide sequence given in the specification), are new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are included for the following:

- (1) a method (I) for diagnosing the presence of a gynecological or testicular cancer in a patient, comprising:
- (i) measuring the levels of CSG in cells, tissues or bodily fluids of the patient; and
- (ii) comparing the measured levels of CSG with the levels found in a normal human control (a change in the measured level of CSG is associated with the presence of the cancer);
- (2) a method (II) for diagnosing and monitoring metastases of a gynecological or testicular cancer, comprising:
- (i) identifying a patient suffering from a cancer that is not known to have metastasized;
- (ii) periodically measuring CSG levels in samples of cells, tissues or fluids from the patient; and
- (iii) comparing the measured levels of CSG with the levels found in a normal human control (an increase in the measured level of CSG is associated with the presence of a cancer that has metastasized);
- (3) a method (III) of staging a gynecological or testicular cancer, comprising:
 - (i) identifying a patient with the cancer;
- (ii) periodically measuring levels of CSG in samples of cells tissues or fluids from the patient; and
- (iii) comparing the measured levels of **CSG** with the levels found in a normal human control (an increase in the measured level of **CSG** is associated with the progression of the cancer and a decrease in the levels is associated with the remission of the cancer);
 - (4) an antibody (Ab) against CSG;
- (5) a method (IV) of imaging a gynecological or testicular cancer comprising administering Ab; and
- (6) a method (V) of treating a gynecological or testicular cancer comprising administering Ab.
 - USE (I) (IV) may used for be diagnosing, staging, imaging

and treating gynecologic and testicular cancers.

ADVANTAGE - Early diagnosis of cancers improves the success rate of therapeutic protocols.

Dwg.0/0

L4 ANSWER 9 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-256657 [22] WPIDS

DOC. NO. CPI: C2000-078328

TITLE: Diagnosing, staging, monitoring, imaging and

treating cancer especially gynecological cancers e.g. breast, ovarian cancer and lung cancer, involves measuring cancer specific gene levels in

cells and body fluids.

DERWENT CLASS: B04 D16

INVENTOR(S): CAFFERKEY, R; RECIPON, H; SALCEDA, S; SUN, Y

PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS INC; (DIAD-N) DIADEXUS LLC

COUNTRY COUNT: 22

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000012758 A1 20000309 (200022)* EN 58

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1109937 A1 20010627 (200137) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO KIND | APPLICATION | DATE |
|------------------|------------------|----------|
| | | |
| WO 2000012758 A1 | .WO 1999-US19655 | 19990901 |
| EP 1109937 A1 | EP 1999-946662 | 19990901 |
| | WO 1999-US19655 | 19990901 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|--------------|
| | | |
| EP 1109937 | Al Based on | WO 200012758 |

PRIORITY APPLN. INFO: US 1998-98880P 19980902

AN 2000-256657 [22] WPIDS

AB WO 200012758 A UPAB: 20000508

NOVELTY - Detecting, diagnosing **metastasis** and staging cancer by measuring levels of cancer specific genes (**CSG**) in cells, tissues or body fluids, is new. Their remission and progression, decreases and increases in **CSG** levels, is also monitored, by periodic sample analysis.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for an antibody (I) against **CSG** which comprises a 2587, 576, 2070, 890, 1709, 406, 2479, 462 or 272 base pair sequence, all fully defined in the specification.

ACTIVITY - Cytostatic. No supporting data given.

MECHANISM OF ACTION - None given.

USE - The methods are useful for detecting, diagnosing, monitoring, staging, prognosing cancers, especially gynecologic cancers which include ovarian, breast, endometrial and uterine

cancer (claimed) and lung cancer. (I) labeled with paramagnetic ions or a radioisotope is useful for imaging cancer and (I) conjugated with a cytotoxic agent is useful for treating cancer (claimed).

ADVANTAGE - The discrimination between metastasized and non-metastasized cancers, which was not possible using prior techniques, can be achieved using this method. Dwg.0/0

L4 ANSWER 10 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-205579 [18] WPIDS

DOC. NO. NON-CPI: N2000-152973 DOC. NO. CPI: C2000-063380

TITLE: Novel methods for diagnosing, monitoring, staging,

imaging and treating colon cancer by measuring the level of colon

specific gene markers.

DERWENT CLASS: B04 D16 S03

INVENTOR(S): MACINA, R A; RECIPON, H; SUN, Y

PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS LLC

COUNTRY COUNT: 22

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000007632 A1 20000217 (200018)* EN 42

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1107798 A1 20010620 (200135) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO K | IND | API | PLICATION | DATE |
|-----------------------------|----------|-----|---|----------|
| WO 2000007632 EP 1107798 | A1 A1 | EP | 1999-US16357 1999-937328 1999-US16357 | 19990720 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|--------------|
| | | |
| EP 1107798 | Al Based on | WO 200007632 |

PRIORITY APPLN. INFO: US 1998-95231P 19980804

AN 2000-205579 [18] WPIDS

AB WO 200007632 A UPAB: 20000412

NOVELTY - A novel method for diagnosing the presence of

colon cancer in a patient comprises measuring

levels of colon specific gene markers

(CSG) in cells, tissues or bodily fluids, and comparing the measured levels of CSG with levels of CSG

from a normal human control, where an increase in measured CSG levels in the patient versus control is associated with

the presence of colon cancer.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) a method of diagnosing metastatic colon

cancer in a patient, comprising:

- (a) identifying a patient having colon cancer that is not known to have metastasized;
- (b) measuring levels of CSG in cells, tissues or bodily fluids in the patient; and
- (c) comparing the measured levels of CSG with levels of CSG from a normal human control, where an increase in measured CSG levels in the patient versus control is associated with a cancer which has metastasized;
- (2) a method of staging colon cancer in a patient, comprising:
 - (a) identifying a patient with colon cancer
- (b) measuring CSG levels in a cell, tissue or bodily fluid sample; and
- (c) comparing levels to a normal human control sample, where an increase in CSG levels is associated with a cancer which is progressing, and a decrease in CSG levels is associated with a cancer which is regressing or in remission;
- (3) a method of monitoring colon cancer in a patient for the onset of metastasis, comprising:
- (a) identifying a patient having colon cancer that is not known to have metastasized;
- (b) periodically measuring CSG levels in a cell, tissue or bodily fluid sample; and
- (c) comparing the levels with a sample obtained from a normal human control where an increase in any one of the periodically measured levels is associated with a cancer that has metastasized;
- (4) a method of monitoring changes in a stage of **colon** cancer in patient, comprising:
 - (a) identifying a patient having colon cancer
- (b) periodically measuring CSG levels in a cell, tissue or bodily fluid sample; and
- (c) comparing levels with a sample obtained from a normal human control, where an increase in any one of the periodically measured levels is associated with a cancer which is progressing in stage and a decrease in any one of the periodically measured levels is associated with a cancer which is regressing in stage or in remission;
- (5) an antibody against a **CSG** which comprises the 1710, 1109 or 1141 base pair (bp) sequence, all fully defined in the specification;
- (6) a method of imaging colon cancer in a patient, comprising administering to the patient the antibody of (5); and
- (7) a method of treating colon cancer in a patient, comprising administering to the patient the antibody of (5).

ACTIVITY - Cytostatic.

MECHANISM OF ACTION - Antibodies to colon specific genes are administered alone or conjugated to cytotoxic agents.

USE - The method is used to detect, monitor, stage or give a prognosis for colon cancer (claimed). The antibodies are used for detection or image localization of the colon specific genes (CSGs).

The antibodies can be conjugated to cytotoxic agent or drug and used to treat colon cancer (claimed).

ADVANTAGE - The methods of the invention are more accurate than prior art clinical methods for staging colon cancer, because they measure colon specific markers, and, unlike pathological staging methods, do not depend on an invasive procedure.

L4 ANSWER 11 OF 24 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.

ACCESSION NUMBER:

Dwg.0/0

2000349854 EMBASE

TITLE:

Flow cytometric analysis of apoptosis and

proliferation in gastric cancer and precancerous

lesion.

AUTHOR:

Yu Qing Guo; Zhao Hua Zhu; Jin Fang Li

CORPORATE SOURCE:

Dr. Y.Q. Guo, Department of Gastroenterology, Baoan

District Hospital of Shenzhen, Shenzhen 518101,

Guangdong Province, China

SOURCE:

World Chinese Journal of Digestology, (2000) 8/9

(983 - 987).

Refs: 54

ISSN: 1009-3079 CODEN: SHXZF2

COUNTRY:

China

DOCUMENT TYPE:

Journal; Article 016 Cancer

FILE SEGMENT:

048 Gastroenterology

LANGUAGE:

Chinese

SUMMARY LANGUAGE: English; Chinese

AIM: To investigate the changes and the possible role of the cell apeptosis index (Al), proliferation index (PI), the ratio of AI to PI and the DNA index (DI) in various gastric diseases. METHODS: The gastric mucosal biopsies taken by endoscopy from 15 cases of chronic superficial gastritis (CSG), 15 cases of chronic atrofic gastritis (CAG), 16 cases of gastritis epithelial dysplasis (Dys) and the specimens of gastric cancer (GC) tissue from surgically resected stomach in 50 patients with gastric cancer were collected in this study. All specimens were fixed in 10% formalin and embedded in paraffin. The apoptosis cells were labelled by TUNEL technic, the AI was expressed as the percentage of positive TUNEL staining cells. The total cell DNA was labelled by propidium index. The PI was expressed as the percentage of S + G2 M phases cells. The AI and PI ratio of AI to PI and the DI were measured by flow cytometry. RESULTS: The AI was significantly lower in GC (6.6%) than that in CAG (11.2%) and Dys (18.3%, P<0.05), but not statistically lower than the AI in CSG (10.2%, P>0.05). The PI in CSG, CAG, Dys and GC were 14.9%, 20.1%, 24.6% and 31.8% respectively. A statistical difference in PI was found between any two groups (P<0.05). The ratio of AI to PI was the lowest in the GC (0.22)compared to CSG (0.65%), CAG (0.57) and Dys (0.72%, P<0.05). The values of PI was significantly related to the depth of tumor invasion, lymph nodes or distant metastasis, tumor stages as well as the type of DNA ploid (P<0.05). All of CSG and CAG were diaploid, however, the aneuploid was found in 25% cases of Dys (4/16) and 82% cases of GC (41/50). The gastric cancers with aneuploid had a significantly higher PI (33.6%) and lower ratio of AI to PI (0.19%) when compared with that in diaploid tumor (PI = 23.4%, AI/PI = 0.35), P<0.05. There was a positive correlation between the AI and PI in both CSG and CAG (r = 0.52 and r

= 0.55, repectively), P<0.05, but such correlation was not seen in the Dys and GC. CONCLUSION: The breakdown of the balance between the cell proliferation and apoptosis gradually developed from CSG to GC. The character of cell kinetics in gastric cancer was the superiority of proliferation to apoptosis due to the increase in proliferation and the decrease in apoptosis. The aneuploid DNA was detected earliest in Dys and became preponderant in GC. The gastric cancer with aneuploid DNA had significantly higher PI and lower ratio of AI to PI. These results suggested that the detection of aneuploid may be useful for a early diagnosis of precarcerous lesion as well as gastric cancer, and for evaluating the malignant degree of tumor.

L4 ANSWER 12 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER: 2000-126383 [11] WPIDS

DOC. NO. NON-CPI: N2000-095292 DOC. NO. CPI: C2000-038417

TITLE: Diagnosing, monitoring and staging colon

cancer.

DERWENT CLASS: B04 D16 J04 S03

INVENTOR(S): MACINA, R A; SUN, Y; YANG, F

PATENT ASSIGNEE(S): (DIAD-N) DIADEXUS LLC

COUNTRY COUNT: 22

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 9960161 A1 19991125 (200011) * EN 29

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: CA JP US

EP 1080227 A1 20010307 (200114) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|------------|------|-----------------|------|
| WO 9960161 | A1 | WO 1999-US10498 | |
| EP 1080227 | A1 | EP 1999-924210 | |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|------------|-------------|------------|
| | | |
| EP 1080227 | Al Based on | WO 9960161 |

PRIORITY APPLN. INFO: US 1998-86266 19980521

AN 2000-126383 [11] WPIDS

AB WO 9960161 A UPAB: 20000301

NOVELTY - Diagnosing the presence, or metastasis, of

colon cancer in a patient, comprising measuring

Colon Specific Gene (CSG)

levels in a cell, tissue or bodily fluid sample of the patient and a control, where increased CSG levels in the patient compared to the control is associated with the presence, or

metastasis, of colon cancer, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for

the following:

- (1) staging colon cancer in a patient, comprising identifying a patient with colon cancer , measuring CSG levels in a cell, tissue or bodily fluid sample and comparing levels to a control sample, where increasing CSG levels is associated with a cancer which is progressing, and decreased levels are associated with a cancer which is regressing or in remission;
- (2) monitoring colon cancer in a patient for the onset of metastasis, comprising identifying a patient having colon cancer that is not known to have metastasized, periodically measuring CSG levels in a cell, tissue or bodily fluid sample, and comparing the levels with a sample obtained from a control where an increase in any one of the periodically measured levels is associated with a cancer that has metastasized; and
- (3) monitoring changes in a stage of colon cancer in patient, comprising identifying a patient having colon cancer, periodically measuring CSG levels in a cell, tissue or bodily fluid sample, and comparing levels with a sample obtained from a control, where an increase in any one of the periodically measured levels is associated with a cancer which is in progressing stage and a decrease in any one of the periodically measured levels is associated with a cancer which is regressing in stage or in remission.

USE - The novel method is used to detect, monitor, stage and give a prognosis for colon cancer.

ADVANTAGE - The invention is more accurate than prior art clinical methods for staging colon cancer, and unlike pathological staging methods, does not depend on an invasive procedure. Dwg.0/0

ANSWER 13 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD L4

ACCESSION NUMBER: 1999-130432 [11]

2000-464055 [38]

CROSS REFERENCE:

DOC. NO. CPI:

C1999-038062

TITLE:

Isolated human colon specific

gene - used to develop products for the diagnosis and treatment of disorders of the

WPIDS

colon, e.g. colon cancer

and metastases.

DERWENT CLASS:

B04 D16

INVENTOR(S): PATENT ASSIGNEE(S): DILLON, P J; LI, Y; SOPPET, D R (HUMA-N) HUMAN GENOME SCI INC

COUNTRY COUNT:

PATENT INFORMATION:

WEEK PG PATENT NO KIND DATE LA US 5861494 A 19990119 (199911)*

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|------------|------|----------------|----------|
| | | | |
| US 5861494 | A | US 1995-468413 | 19950606 |

Shears 308-4994 Searcher :

PRIORITY APPLN. INFO: US 1995-468413 19950606

AN 1999-130432 [11] WPIDS

CR 2000-464055 [38]

AB US 5861494 A UPAB: 20000823

- (A) An isolated polynucleotide (PN) which comprises a member selected from:
- (a) a PN sequence encoding a polypeptide comprising amino acids 2 to 158 of a 158 amino acid sequence (II) as given in the specification, and
 - (b) the full complement of (a).

Also claimed are:

- (1) a recombinant vector comprising a PN as in (A), where the PN is DNA;
- (2) a recombinant host cell comprising a PN as in (A), where the PN is DNA;
 - (3) an isolated PN comprising a member selected from:
- (a) a PN sequence encoding the same mature polypeptide encoded by a human cDNA in ATCC No. 97129, and
 - (b) the full complement of (a);
- (4) an isolated PN comprising a PN sequence that will hybridise under stringent conditions to a member selected from (a) and (b) as in (A);
- (5) an isolated PN comprising a PN sequence that will hybridise under stringent conditions with a member selected from (a) and (b) as in (4);
- (6) a method of making a recombinant vector comprising inserting an isolated PN as in (3), (4) or (5) into a recombinant vector, where the PN is DNA, and
- (7) a recombinant host cell comprising a PN as in (3), (4) or (5), where the PN is DNA.

USE - The PNs, which represent a human colon specific gene can be used to develop products for the diagnosis of a disorder of the colon, e.g. colon cancer or metastases. The products can also be used to screen for agonists or antagonists for the polypeptides.

The antagonists may be used to treat **colon cancer**, since they interact with the function of **colon** specific polypeptides in a manner to inhibit natural function which is necessary for the viability of **colon cancer** cells. The products can also be used for the production of antibodies and for the identification of receptors for the polypeptides.

Dwg.0/1

L4 ANSWER 14 OF 24 MEDLINE DUPLICATE 1

ACCESSION NUMBER: 1999243161 MEDLINE

DOCUMENT NUMBER: 99243161 PubMed ID: 10226574

TITLE: Extract of Solanum muricatum (Pepino/CSG)

inhibits tumor growth by inducing apoptosis.

AUTHOR: Ren W; Tang D G

CORPORATE SOURCE: Virotech Canada Inc., Windsor, ON, Canada..

wpren@mnsi.net

SOURCE: ANTICANCER RESEARCH, (1999 Jan-Feb) 19 (1A) 403-8.

Journal code: 59L; 8102988. ISSN: 0250-7005.

PUB. COUNTRY: Greece

Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

Priority Journals FILE SEGMENT:

199905 ENTRY MONTH:

Entered STN: 19990601 ENTRY DATE:

> Last Updated on STN: 19990601 Entered Medline: 19990520

BACKGROUND: Apoptosis, or programmed cell death, is characterized by AΒ certain distinct morphological and biochemical features. Most chemotherapeutic drugs exert their anti-tumor effects by inducing apoptosis. Therefore, an effective compound inducing apoptosis appears to be a relevant strategy to suppress various human tumors. In a search for tumor inhibitors from various kinds of plants, we found that extracts from Solanum muricatum (CSG) can inhibit tumor growth both in vivo and in vitro by inducing apoptosis. MATERIALS AND METHODS: A lyophilized aqueous fraction extracted from Solanum muricatum (CSG4) was used in this study. The human cell lines tested include: prostate (PC3, DU145), stomach (MKN45), liver (QGY-7721, SK-HEP-1), breast (MDA-MB-435), ovarian (OVCAR), colon (HT29) and lung (NCI-H209) cancer cells; NHP (prostate), HUVEC (umbilical vein endothelial cell), and WI-38 (lung diploid fibroblasts) normal cells. The cell survival was determined by either Cell Titer MTS cell proliferation kit or trypan blue dye exclusion assay. The apoptosis was analyzed by (a) apoptotic morphology by light microscopy; (b) DNA ladder formation; (c) PARP cleavage assay. RESULTS: a) CSG possesses selective cytotoxic activity against all the tumor cell lines being tested. The LD50 value is 561-825 micrograms/ml. b) CSG showed a much lower cytotoxicity to NHP, HUVEC and WI-38 normal cell lines with LD50 value being 2.8-3.2 mg/ml, which is 3-6 fold higher than on tumor cells. c) The in vivo study demonstrated that injection of CSG (100 micrograms) directly into tumor mass can reduce the tumor volume dramatically in nude mice inoculated with MKN45 gastric cancer cells. d) CSG-mediated tumor growth inhibition is through induction of apoptotic cell death, as manifested by (a) typical apoptotic morphology; (b) DNA ladder formation; and (c) PARP cleavage assay. CONCLUSION: Taken together, the present study suggests, for the first time, that CSG may represent promising new chemical entity which preferentially targets various tumor cells by triggering apoptosis.

DERWENT INFORMATION LTD ANSWER 15 OF 24 WPIDS COPYRIGHT 2002 WPIDS

ACCESSION NUMBER: 1998-229823 [20]

C1998-071736 DOC. NO. CPI:

TITLE: Colon-specific nucleic acids - useful as

probes for detecting colon cancer

micrometastases.

B04 D16 DERWENT CLASS:

INVENTOR(S): ROSEN, C; YU, G

(HUMA-N) HUMAN GENOME SCI INC PATENT ASSIGNEE(S):

COUNTRY COUNT:

PATENT INFORMATION:

PG PATENT NO KIND DATE WEEK LA US 5733748 50 A 19980331 (199820)*

> Shears 308-4994 Searcher :

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|------------|------|----------------|----------|
| | | | |
| US 5733748 | A | US 1995-469667 | 19950606 |

PRIORITY APPLN. INFO: US 1995-469667 19950606

AN 1998-229823 [20] WPIDS

AB US 5733748 A UPAB: 19980520

A new isolated polynucleotide (I) comprises a sequence at least 95% identical to a sequences encoding polypeptides that are either: (a) a 167 amino acid (aa) sequence; (b) aa 2-135 of a 135 aa sequence; (c) a 228 aa sequence; (d) a 163 aa sequence; (e) an 81 aa sequence; (f) aa 2-323 of a 323 aa sequence; (g) a 156 aa sequence; or (h) the complements of sequences as in (a)-(g).

Also claimed are: (1) a recombinant vector comprising (I); (2) a recombinant host cell containing (1); and (3) an isolated polynucleotide comprising a sequence at least 95% identical to a sequence encoding a mature polypeptide encoded by the human cDNA in ATCC 97102 or its complement.

USE - The polynucleotides are partial or full-length cDNA clones of colon-specific genes and can be used as probes to detect expression of the corresponding human genes, e.g. in diagnostic assays for detecting micrometastases of colon cancer. The recombinant cells can be used to produce the polypeptides, in order that antibodies can be raised and used in further screening or diagnostics.

Dwg.0/13

L4 ANSWER 16 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD

ACCESSION NUMBER:

1997-043162 [04] WPIDS

DOC. NO. NON-CPI:

N1997-035728

DOC. NO. CPI:

C1997-013821

TITLE:

New isolated colon specific

gene - used to develop prods. for use in the diagnosis and treatment of colon

disorders, partic. colon cancer

DERWENT CLASS:

B04 D16 S03

INVENTOR(S):

DILLON, P J; LI, Y; SOPPET, D R (HUMA-N) HUMAN GENOME SCI INC

PATENT ASSIGNEE(S):

60

COUNTRY COUNT:
PATENT INFORMATION:

| PATENT | NO | KIND | DATE | WEEK | LA | PG |
|--------|----|------|------|------|----|----|
| | | | | | | |

WO 9639541 A1 19961212 (199704)* EN 64

RW: AT BE CH DE DK ES FR GB GR IE IT KE LU MC MW NL OA PT SD SE SZ UG

W: AM AT AU BB BG BR BY CA CH CN CZ DE DK ES FI GB GE HU JP KE KG KP KR KZ LK LT LU LV MD MG MN MW MX NO NZ PL PT RO RU SD

SE SI SK TJ TT UA US UZ VN

AU 9528180 A 19961224 (199715) EP 833948 A1 19980408 (199818) EN

R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

CN 1194009 A 19980923 (199906)

JP 11506920 W 19990622 (199935) 57

AU 711346 B 19991014 (200001) KR 99022532 A 19990325 (200023)

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|-------------|------|----------------|----------|
| WO 9639541 | A1 | WO 1995-US7169 | 19950606 |
| AU 9528180 | A | AU 1995-28180 | 19950606 |
| | | WO 1995-US7169 | 19950606 |
| EP 833948 | A1 | EP 1995-923729 | 19950606 |
| | | WO 1995-US7169 | 19950606 |
| CN 1194009 | A | CN 1995-197931 | 19950606 |
| | | WO 1995-US7169 | 19950606 |
| JP 11506920 | W | WO 1995-US7169 | 19950606 |
| | | JP 1997-500365 | 19950606 |
| AU 711346 | В | AU 1995-28180 | 19950606 |
| | | WO 1995-US7169 | 19950606 |
| KR 99022532 | A | WO 1995-US7169 | 19950606 |
| | | KR 1997-709013 | 19971206 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|-------------|----------------|---------------|
| AU 9528180 | A Based on | WO 9639541 |
| EP 833948 | Al Based on | WO 9639541 |
| JP 11506920 | W Based on | WO 9639541 |
| AU 711346 | B Previous Pub | 1. AU 9528180 |
| | Based on | WO 9639541 |
| KR 99022532 | A Based on | WO 9639541 |

PRIORITY APPLN. INFO: WO 1995-US7169 19950606

AN 1997-043162 [04] WPIDS

AB WO 9639541 A UPAB: 19970122

An isolated polynucleotide (PN) comprises a member selected from:
(a) a PN encoding the polypeptide comprising amino acids 1-158 of a
158 amino acid sequence given in the specification; (b) a PN which
encodes a mature polypeptide encoded by the DNA contained in ATCC
Deposit No. 97129; (c) a PN capable of hybridising to and which is
at least 70% identical to a PN of (a) or (b); and (d) a PN fragment
of a PN of (a), (b) or (c).

USE - The PNs can be used for the diagnosis of disorders of the colon in hosts. The polypeptide and its (ant)agonists can be used for the treatment of disorders of the colon, partic. colon cancer.

Dwg.0/1

L4 ANSWER 17 OF 24 WPIDS COPYRIGHT 2002 DERWENT INFORMATION LTD ACCESSION NUMBER: 1997-043054 [04] WPIDS

DOC. NO. CPI:

C1997-013713

TITLE:

Human colon specific

genes and their expression products detection of which, in non-colon tissue
samples, can be used as indication of colon

cancer metastasis.

DERWENT CLASS:

B04 D16

INVENTOR(S): ROSEN, C A; YU, G

PATENT ASSIGNEE(S): (HUMA-N) HUMAN GENOME SCI INC

COUNTRY COUNT:

60

PATENT INFORMATION:

| PA: | FENT | NO | 1 | KINI | D D | ATE | | W | EEK | |] | LA | P | 3 | | | | | | | |
|-----|------|------|-------|-------|------|------|------|------|-------------------|------|-----------|--------|------------|-----------------|----|----|----|----|----|----|----|
| WO. | 963 | 941 | 9 | Α | 1 1 | 996 | 1212 | 2 (| 199 | 704° |) * 1 | EN | 88 | - - 3 | | | | | | | |
| | RW: | | | | | | | | | | | | | | MC | MW | NL | OA | PT | SD | SE |
| | | | UG | | | | | | | | | | | | | | | | | | |
| | W: | AM | AT | ΑU | BB | ВG | BR | BY | CA | CH | CN | CZ | DΕ | DK | ES | FI | GB | GE | HU | JΡ | KE |
| | | KG | KP | KR | ΚZ | LK | LT | LU | $rac{r}{\Lambda}$ | MD | MG | MN | MW | MX | ИО | ΝZ | PL | PT | RO | RU | SD |
| | | SE | ŞΙ | SK | ТJ | TT | UA | US | UZ | VN | | | | | | | | | | | |
| ΑU | 952 | 820 | 5 | Α | 19 | 996: | 1224 | 1 (: | 199 | 715 |) | | | | | | | | | | |
| EΡ | 847 | 398 | | A1 | 1 19 | 9980 | 0617 | 7 (: | 199 | 828 |) I | ΞN | | | | | | | | | |
| | R: | ΑT | ΒE | CH | DE | DK | ES | FR | GB | GR | ΙE | IT | $_{ m LI}$ | LU | MC | NL | PT | SE | | | |
| JP | 1150 | 0634 | 12 | W | 19 | 9990 | 0608 | 3 (: | 199 | 933 |) | | 7: | L | | | | | | | |

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE | | |
|-------------|------|----------------|----------|--|--|
| WO 9639419 | A1 | WO 1995-US7289 | 19950606 | | |
| AU 9528205 | A | AU 1995-28205 | 19950606 | | |
| | | WO 1995-US7289 | 19950606 | | |
| EP 847398 | A1 | EP 1995-923764 | 19950606 | | |
| | | WO 1995-US7289 | 19950606 | | |
| JP 11506342 | W | WO 1995-US7289 | 19950606 | | |
| | | JP 1997-500380 | 19950606 | | |

FILING DETAILS:

| PAT | TENT NO | KIND | | | PAT | ENT NO | |
|-----|----------|------|-------|----|-----|---------|--|
| AU | 9528205 | A | Based | on | WO | 9639419 | |
| EP | 847398 | A1 | Based | on | WO | 9639419 | |
| JΡ | 11506342 | W | Based | on | WO | 9639419 | |

PRIORITY APPLN. INFO: WO 1995-US7289 19950606

AN 1997-043054 [04] WPIDS

AB WO 9639419 A UPAB: 19970122

A novel isolated polynucleotide (I), is selected from; (a) a polynucleotide encoding the same polypeptide as a polynucleotide having a 1129 bp nucleic acid sequence given in the specification, or an at least 70% identical hybrid; or (b) a polynucleotide encoding the same mature polypeptides as a human gene having a coding portion, which includes DNA having at least 90% identity to the DNA one of nine nucleic acid sequences given in the specification, which represent fragments of colon specific genes, or a DNA included in ATCC 97102.

USE - The novel isolated polynucleotide, comprises 1 of 13 human colon specific genes, designated CSG1-CSG13, which are primarily expressed in colon derived tissues. Transcription of these human genes in a non-colon tissue sample can be used as an indication of a colon disorder (i.e. colon cancer metastases); specifically the detection of an altered level of RNA transcribed from one of the human genes, DNA complementary to the RNA or an expression prod. (e.g. detected in an immunoassay using the

antibody) (claimed). The polypeptide and cpd. (which may be a polypeptide expressed in vivo via the admin. of a polynucleotide encoding the cpd.) can be used for the treatment of a patient in need of CSG protein or CSG protein inhibition, respectively (claimed), e.g. a colon cancer patient.

Dwg.0/13

L4 ANSWER 18 OF 24 CANCERLIT

ACCESSION NUMBER: 96647047 CANCERLIT

DOCUMENT NUMBER: 96647047

TITLE: Phase II first line chemotherapy (CT) study with

docetaxel (taxotere) and prophylactic premedication of fluid retention (FR) in patients (pts) with

metastatic (MTS) or locally advanced breast cancer (ABC): EORTC Clinical Screening Group (

CSG) (Meeting abstract).

AUTHOR: Fumoleau P; Krakowski I; Chevallier B; Roche H;

Kerbrat P; Dieras V; Azli N; Rios M; Riva A; Lentz M

A; van Glabbeke M

CORPORATE SOURCE: Centre R. Gauducheau, Nantes, France.

SOURCE: Non-serial, (1995). EORTC Early Drug Development

Meeting 1995, June 21-24, 1995, Corfu, Greece.

DOCUMENT TYPE: (MEETING ABSTRACTS)

(CLINICAL TRIAL, PHASE II)

(CLINICAL TRIAL)

FILE SEGMENT: ICDB LANGUAGE: English

199607 ENTRY MONTH: CSG has already reported on the activity and the toxicity of Docetaxel as first line CT in pts with mts or locally ABC (ASCO A115, 1994). This multi-center study was performed in order to confirm efficacy and to evaluate efficacy of prophylactic premedication including dexchlorpheniramine iv 5 mg and ranitidine iv 50 mg 30 mn before CT plus prednisolone po 130 mg 12 and 6 hr before CT in order to reduce the incidence and severity of FR observed in previous studies. From 08/93 to 05/94 37 pts were included and all were evaluable for response and safety. Pts: median age = 48(29-65); Ps who at baseline was PS = 0(48.6%), PS = 1(43.2%), PS = 2(8.1%); Nb of metastatic sites was 1(21.6%), 2(29.7%), greater than 2(48.6%); Mts sites: liver (40.5%), lung (37.8%), bone (51.4%), lymph nodes (48.6%), skin (18.9%), breast (18.9%); 24 pts received prior neoadjuvant and/or adjuvant CT with anthracyclines in 87.5%; Median time between last CT and Docetaxel was 32.1 (2.8-143 months). All responses were reviewed by the same independent board. Treatment:total number of cycles= 200; median number of cycles = 5 (1-10); median cumulative dose= 499 (97.6-994 mg/m2); median dose intensity = 32.7 (19.6-33.8 mg/m2/w). Results (NCI-CTC criteria): 2 CR, 23 PR, 8 NC, 4 PD, giving a RR of 67.6% (95% CI:50.2-82%). Median duration of responses not reached (9+-36+w); median time to response= 7+(1+-22+w); median time to progression on 31/05/94 = 31+(1-36+w). Response by site was skin 100%, lymph nodes 78.6%, liver 76.9%, breast 66.7%, lung 0%. RR was not affected by prior or no CT nor nb of organs involved (1 vs 2 vs

greater than 2). Safety: overall grade 4 neutropenia occurred in 89% with median duration of 6 days (2-14) and febrile neutropenia in 35.1%. Other grade 3-4 NCI toxicities included alopecia (100%), stomatitis (5.4%) and skin (2.7%). HSR toxicity and neurotoxicity

were only grade 1-2 and occurred, respectively, in 16.2 and 81.1% of pts. Nails disorders occurred in 64.9% and was moderate in 13.5%; asthenia in 73% and was moderate in 21.6%. FR was a reason for study discontinuation in 43.2%. Median cumulative dose to onset of FR was 301 mg/m2 (98-595) and to treatment discontinuation due to retention was 698 mg/m2 (98-995+). This syndrome was slowly reversible. Conclusions: this study confirms the overall activity of Docetaxel particularly in liver mts pts. The main acute toxicities observed are easy to handle. The premedication proposed failed to reduce the incidence and/or to delay the onset of fluid retention.

L4 ANSWER 19 OF 24 CANCERLIT

ACCESSION NUMBER: 95613243 CANCERLIT

DOCUMENT NUMBER: 95613243

TITLE: Phase II first line chemotherapy (CT) study with

docetaxel (Taxotere) and prophylactic premedication of fluid retention (FR) in patients (pts) with metastatic (mts) or locally advanced breast cancer (ABC). EORTC clinical screening group (

CSG) (Meeting abstract).

AUTHOR: Krakowski I; Rios M; Fumoleau P; Chevaller B; Roche

H; Kerbrat P; Deras V; Azli N; Bougon N; Riva A; et

al

CORPORATE SOURCE: Centre A. Vautrin, CRLCC, 54511 Vandoeuvre les Nancy,

France.

SOURCE: Proc Annu Meet Am Soc Clin Oncol, (1995). Vol. 14,

pp. A87.

ISSN: 0732-183X. (MEETING ABSTRACTS)

(CLINICAL TRIAL)
(MULTICENTER STUDY)

(CLINICAL TRIAL, PHASE II)

FILE SEGMENT: ICDB LANGUAGE: English

DOCUMENT TYPE:

199511 ENTRY MONTH: CSG has already reported on the activity and the toxicity of docetaxel as first line CT in pts with mts or locally ABC (ASCO 94 A115). This multicenter study was performed in order to confirm efficacy and to evaluate efficacy of prophylactic premedication including dexchlorpheniramine iv 5 mg and ranitidine iv 50 mg 30 min before CT plus prednisolone po 130 mg 12 and 6 hr before CT in order to reduce the incidence and severity of FR observed in previous studies. From 08/93 to 05/94, 37 pts were included and all were evaluable for response and safety. Pts: median age=48 (29-65), PS WHO at baseline was PS=0 (48.6%), PS=1 (43.2%), PS=2 (8.1%); metastatic sites: 1 (21.6%), 2 (29.7%), greater than 2 (48.6%) (visceral involvement in 75.7%); mts locale: liver (40.5%), lung (37.8%), bone (51.4%), lymph nodes (48.6%), skin (18.9%), breast (18.9%); 24 pts (64.9%) received prior neoadjuvant and/or adjuvant CT with anthracycline in 87.5%; median time between last CT and docetaxel was 32.1 (12.8-143.0) months. All responses were reviewed by the same independent board. Treatment: total number of cycles=200, median 5 (1-10); median cumulative dose=499 (97.6-994.5 mg/m2); median dose intensity=32.7 (19.6-33.8 mg/m2/w). Results (NCI-CTC criteria): 2 CR, 23 PR, 8 NC, 4 PD, RR was 67.6% (95% CI: 50.2-82%); median duration of response not reached (9+ to 36+ w); median time to response=7+ (1+ to 22+ w); median time to progression on 31/05/94=31+ (1-36+ w). Response by site: skin 100%, lymph nodes

78.6%, liver 76.9%, breast 66.7%, lung 0%. RR is not affected by prior or no CT nor number of organs involved (1 vs 2 vs greater than 2). Safety: overall grade 4 neutropenia occurred in 89.0% with median duration of 6 days (2-14) and febrile neutropenia in 35.1%. Other grade 3-4 NCI toxicities included alopecia (100%), stomatitis (5.4%) and skin (2.7%). HSR toxicity and neurotoxicity were only Grade 1-2 and occurred, respectively, in 16.2% and 81.1% of pts. Nails disorder occurred in 64.9% and was moderated in 13.5%; asthenia in 73% and was moderated in 21.6%. FR occurred in 89.2% and was moderated in 32.4% or severe in 10.8%. FR was a reason for study discontinuation in 43.2%. Median cumulative dose to onset of FR was 301 mg/m2 (98+ to 595) and to treatment discontinuation due to retention was 698 mg/m2 (98+ to 995). This syndrome was slowly reversible. Conclusions: This study confirmed the overall activity of docetaxel particularly with liver mts. The main acute toxicities observed were easy to handle. The premedication proposed failed to reduce the incidence and/or to delay the onset of fluid retention. (C) American Society of Clinical Oncology 1997

ANSWER 20 OF 24 BIOSIS COPYRIGHT 2002 BIOSIS

ACCESSION NUMBER:

1995:15068 BIOSIS

DOCUMENT NUMBER:

PREV199598029368

TITLE:

Activity of Taxotere (Docetaxel) in liver metastasis (Mts) of advanced breast cancer

(ABC): Analysis on 17 patients (pts), experience of the EORTC Clinical Screening Cooperative Group (

AUTHOR(S):

Chevallier, B. (1); Kerbrat, P.; Dieras, V.; Maugard-Louboutin, C.; Roche, H.; Misset, J. L.; Lentz, M. A.; Azli, N.; Klink-Alakl, M.; Fumoleau, P.

CORPORATE SOURCE:

SOURCE:

(1) Centre Henri Becquerel, Rouen France

Breast Cancer Research and Treatment, (1994) Vol. 32,

No. SUPPL., pp. 34.

Meeting Info.: 17th Annual San Antonio Breast Cancer Symposium on Breast Cancer Research and Treatment San

Antonio, Texas, USA December 6-10, 1994

ISSN: 0167-6806.

DOCUMENT TYPE:

LANGUAGE:

Conference English

ANSWER 21 OF 24 CANCERLIT L4

92678222 CANCERLIT ACCESSION NUMBER:

DOCUMENT NUMBER:

92678222

TITLE:

EOI OSTEOSARCOMA TRIALS (MEETING ABSTRACT).

AUTHOR:

Burgers J M; van Glabbeke M; Souhami R; Bramwell V

CORPORATE SOURCE:

SOURCE:

No affiliation given. Med Pediatr Oncol, (1990). Vol. 18, No. 5, pp. 223.

DOCUMENT TYPE:

(CLINICAL TRIAL) (MULTICENTER STUDY)

FILE SEGMENT:

ICDB

LANGUAGE: ENTRY MONTH: English 199201

The European Osteosarcoma Intergroup (EOI) was composed from the SIOP, the EORTC, the MRC and the UKCCSG; now the CSG from Canada is also participating. The main current study 80861 concerns nonmetastatic operable osteosarcoma of the limb. Two chemotherapy schedules are compared with local treatment (loc tr) at week 9. In

> Shears 308-4994 Searcher :

arm 1, 6 courses of cisplatin (CP) 100 mg/m2, and Adriamycin (A) 25 mg/m2 on days 1, 2, 3 are given at 3 weekly intervals with a greater interval at time of surgery. The 2nd arm consists of a modified T10 protocol including high-dose methotrexate (Mt) with a total duration of 42 wk. In several countries Mt is specially available for such study purposes. At March 1, 1990, 231 patients (pts) were registered, 3 quarters coming from the UK. More participation from the continental SIOP members is welcomed. The goal is to reach a total of 400 pts. The previous protocol 80831 collected 207 pts in the neoadjuvant and adjuvant setting from limb osteosarcoma. The same arm 1 as above was compared to 4 courses of CP and A, preceded at 10 days by 8 mg/m2 Mt with leukovorin rescue. At 3 yr disease-free survival (DFS) is 65% and 41% for the 2 and 3 drug arm, respectively, with equal survival 65%. For planned conservative surgery (cons S) DFS = 56% and for amputation 41% with equal survival. The difference in DFS between the 2 drug arms could be demonstrated for cons S. Local recurrence occurred in 9% of pts. Histologic grading is being performed. Current pilot studies: (a) A, CP and ifosfamide (PIA) in metastatic or inoperable cases and loc tr if possible at week 9, study coordinator Dr Voute; (b) A, CP with loc tr at week 9 for nonosteosarcoma spindle cell tumors or bone, study coordinator Dr V Bramwell.

ANSWER 22 OF 24 CANCERLIT

ACCESSION NUMBER: 82610591 CANCERLIT

DOCUMENT NUMBER: 82610591

[TWO CASES OF SEBACEOUS CARCINOMA OBSERVED AT THE S. TITLE:

SPIRITO HOSPITAL OF CASALE MONTEFERRATO IN 1980]. SU DUE CASI DI CARCINOMA SEBACEO OSSERVATI PRESSO L'OSPEDALE 'S. SPIRITO' DI CASALE MONTEFERRATO

NELL'ANNO 1980.

Deregibus P; Battezzati G AUTHOR:

Divisione di Chirurgia Generale, Ospedale 'S. CORPORATE SOURCE:

Spirito', Casale Monteferrato (Alessandria), Italy. Minerva Med, (1982). Vol. 73, No. 5, pp. 213-217.

SOURCE: ISSN: 0026-4806.

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

FILE SEGMENT: ICDB Italian LANGUAGE: 198205 ENTRY MONTH:

Two patients (1 man, 1 woman; 56 and 78 yr old, respectively) with carcinomas of the sebaceous gland (CSG) were reported. In the woman, a cyst had been removed 20 yr earlier at the right side of the face under the hair; this cyst was close to a tumor that started growing 10 yr later, that reached the size of a small mandarine, and that was without local or distant metastases . The man had a perianal tumor that was removed a few months prior to the appearance of a carcinoma of the surrounding skin; metastases were observed in the groin, rib, shoulder, and skull (behind the left eye), but not in the spine. This patient died 2 mo later. (6 Refs)

ANSWER 23 OF 24 CANCERLIT

82610588 CANCERLIT ACCESSION NUMBER:

DOCUMENT NUMBER: 82610588

TITLE: [COMPARATIVE EVALUATION OF CHOLECYSTOGRAPHY,

CHOLANGIOGRAPHY, ECHOTOMOGRAPHY, AND

CHOLESCINTIGRAPHY IN SURGICAL BILE DUCT DISORDERS].

308-4994 Searcher : Shears

VALUTAZIONE COMPARATIVA FRA COLECISTOGRAFIA, COLANGIOGRAFIA, ECOTOMOGRAFIA E COLESCINTIGRAFIA NELLE AFFEZIONI CHIRURGICHE DELLE VIE BILIARI.

AUTHOR: Sgro M; Mure G; Campanoni V; Clerici R

CORPORATE SOURCE: I Divisione Chirurgia Generale, Ospedale Generale

Provinciale di Gallarate, Gallarate, Italy.

SOURCE: Minerva Med, (1982). Vol. 73, No. 3/4, pp. 109-114.

ISSN: 0026-4806.

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

FILE SEGMENT: ICDB
LANGUAGE: Italian
ENTRY MONTH: 198205

Cholecystography (CCG), cholangiography (CAG), echotomography (ETG), and cholescintigraphy (CSG) were comparatively evaluated in 70 patients (27 men, 43 women; 22-76 yr old) with bile duct disease. Of the 70 patients, 30 had cholelithiasis and surgery, 20 biliary cyst, and 20 jaundice (JA). Results of of the patients with cholelithiasis or biliary cysts showed that the ETG diagnosed correctly 27/30 patients with cholelithiasis and 19/20 patients with biliary cyst. Among the patients with the JA, 6 had pancreatic cancer (PA), 4 had cancer of the hepatic biliary duct (CHBD), and 1 had hilar metastases. Patients with JA and bilirubin greater than 4 mg% should be examined with the aid of the ETG and the CSG, since the CAG cannot be used. The ETG diagnosed correctly 4/6 patients with PA and 2/4 patients with CHBD; the CSG diagnosed correctly the 2/6 patients with the PA. Protocols developed and used in diagnosis of the patients with or without JA were presented. (69 Refs)

L4 ANSWER 24 OF 24 CANCERLIT

ACCESSION NUMBER: 79600632 CANCERLIT

DOCUMENT NUMBER: 79600632

TITLE: CARCINOEMBRYONIC ANTIGEN (CEA) AS A MONITOR OF

CRYOSURGICAL TREATMENT OF PATIENTS WITH RECTAL

CARCINOMA.

AUTHOR: Lamerz R; Feifel G; Kohl H J; Lutz H

CORPORATE SOURCE: Medizinischen Klinik II, Klinikum Grosshadern der

Universitat Munchen, Marchioninistrasse 15, 8000

Munchen 70, W. Germany.

SOURCE: Fortschr Med, (1978). Vol. 96, No. 41 2071-2072, pp.

2074-2075.

ISSN: 0015-8178.

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

FILE SEGMENT: ICDB
LANGUAGE: German
ENTRY MONTH: 197901

The serum levels of CEA were studied in 39 patients (31 men, 8 women; 40-84 yr) with histologically proven rectal carcinoma. All these patients were treated with cryosurgery (CSG) using liquid nitrogen at -110 to -160 C. Cryosurgery was used because the tumor was within 15 cm of the rectum, inoperable, or the patient refused other surgery. An I125 radioimmunoassay was used for CEA; 3 nanog/ml was established as the upper level of normal by studies on healthy persons. Of the 23 patients whose tumors were reduced by CSG (group 1), 7 showed a clear decrease in CEA levels after CSG, 10 showed no change, 5 had markedly increasing levels (12-1,853 nanog/ml), and 2 had variable levels between 3-6 nanog/ml. All five patients with marked increases were found to have distant

metastases in the lungs or liver. Of the 11 patients whose tumors progressed after CSG (group 2), 9 showed an increase in CEA (3 had normal CEA levels before surgery) and 2 had CEA levels that remained close to normal (5 nanog/ml or less). Five patients had tumors unchanged by CSG: CEA levels remained in the normal range in two, CEA levels remained elevated without change in two patients, and a small increase in CEA was seen in one terminal patient. In group 1 patients in whom metastases were not found, only 2/18 had an increasing or elevated CEA level after treatment, compared with 9/11 in group 2 (p less than 0.001). In five patients with distant metastases, CEA increases or constantly elevated values were found; this response was found in only 2/18 without metastases (p less than 0.005). Serum CEA levels, determined before and after CSG, seem to be useful indicators of tumor progression and/or distant metastases. (21 Refs)

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